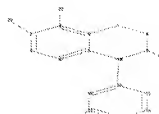
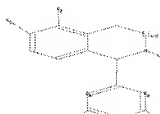


***** Welcome to STN International *****
 ***** STN Columbus *****

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=> Uploading C:\Program Files\Stnexp\Queries\Queries\10591174.str



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20 22 24 29 30
ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16
chain bonds :
2-24 3-22 9-20 10-14 12-30 16-29
ring bonds :
1-2 1-6 2-3 3-4 4-5 4-7 5-6 5-10 7-8 8-9 9-10 11-12 11-16 12-13 13-14
14-15 15-16
exact/norm bonds :
2-24 3-22 4-5 4-7 5-6 5-10 7-8 8-9 9-10 9-20 10-14 11-12 11-16 12-13
12-30 13-14 14-15 15-16 16-29
normalized bonds :
1-2 1-6 2-3 3-4
isolated ring systems :
containing 11 :
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G1:C,H

G2:C,H,O,S,X

G3:C,O,S

G4:C,N

G5:C,O,X

Match level :

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1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 20:CLASS 22:CLASS 24:CLASS
29:CLASS 30:CLASS
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=> s 11 sam

L2 3 SEA SSS SAM L1

=> s 11 full
 L3 574 SEA SSS FUL L1

=> file caplus

=> s 13
 L4 75 L3

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 (PD<20040300)

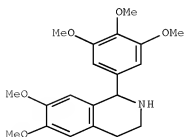
L5 61 L4 AND PD< MARCH 2004

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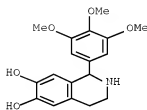
L5 ANSWER 1 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2004:1079594 CAPLUS Full-text
 DN 143:70979
 TI The chromatographic data in QSAR assay of TIQs derivatives with
 β 2-adrenergic activity
 AU Brzezinska, Elzbieta; Stolarska, Justyna
 CS Department of Analytical Chemistry, Medical University of Lodz, Lodz,
 90-151, Pol.
 SO Acta Poloniae Pharmaceutica (2004), 61(4), 249-254
 CODEN: APPHAX; ISSN: 0001-6837
 PB Polish Pharmaceutical Society
 DT Journal
 LA English
 AB We performed QSAR anal. of β 2-adrenergic activity and chromatog. data of
 4,6,8-trihydroxy-, 6,7-dihydroxy- and 6,7-dimethoxy-1,2,3,4-
 tetrahydroisoquinoline derivs. TLC plates (silica gel NP 60 F254 and silica
 gel RP2 60F254 silanised precoated), impregnated with solns. of analogs of the
 selected amino acids were used as β 2-agonistic and antagonistic interaction
 models. QSAR anal. of the β 2-adrenergic activity and the chromatog. data of
 the solutes were made. A correlation between biol. data and behavior of the
 examined compds. in a chromatog. modifiable environment (S1-S3) was
 investigated by the linear regression anal. method.

IT 33033-84-0 188553-85-7
 RL: PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic
 use); BIOL (Biological study); USES (Uses)
 (chromatog. data in QSAR assay of
 6,7-dimethoxy-1,2,3,4-tetrahydroisoquinoline derivs. with
 β 2-adrenergic activity)

RN 33033-84-0 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
 (CA INDEX NAME)



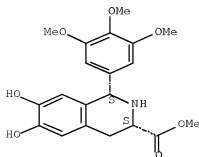
RN 188553-85-7 CAPLUS
 CN 6,7-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



RE.CNT 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

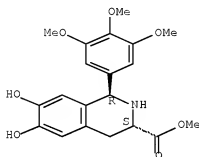
L5 ANSWER 2 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2004:571544 CAPLUS [Full-text](#)
 DN 141:296185
 TI Asymmetric Pictet-Spengler reactions. Synthesis of tetrahydroisoquinoline derivatives from L-DOPA
 AU Wang, Ye; Liu, Zhan Zhu; Chen, Shi Zhi; Liang, Xiao Tian
 CS Institute of Materia Medica, Peking Union Medical College and Chinese Academy of Medical Sciences, Beijing, 100050, Peop. Rep. China
 SO Chinese Chemical Letters (2004), 15(5), 505-507
 CODEN: CCLEE7; ISSN: 1001-8417
 PB Chinese Chemical Society
 DT Journal
 LA English
 OS CASREACT 141:296185
 AB The cis-1-substituted-6,7-dihydroxy-1,2,3,4-tetrahydroisoquinoline-3-carboxylic acid esters can be obtained in a highly diastereoselective fashion through 1,3-induction Pictet-Spengler cyclization of the L-DOPA (3,4-dihydroxyphenylalanine) Me ester with aromatic or aliphatic aldehydes under acidic conditions. Their epimers are also obtained as minor products.
 IT 764660-42-6P 764660-55-1P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (asym. Pictet-Spengler reactions for preparation of tetrahydroisoquinoline derivs. from 3,4-Dihydroxyphenylalanine Me ester and aldehydes)
 RN 764660-42-6 CAPLUS
 CN 3-Isoquinolinecarboxylic acid, 1,2,3,4-tetrahydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, methyl ester, (1R,3R)-rel- (CA INDEX NAME)

Relative stereochemistry.



RN 764660-55-1 CAPLUS
 CN 3-Isoquinolinecarboxylic acid, 1,2,3,4-tetrahydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, methyl ester, (1R,3S)-rel- (CA INDEX NAME)

Relative stereochemistry.



OSC.G 7 THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD (7 CITINGS)
 RE.CNT 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 3 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2004:362287 CAPLUS [Full-text](#)

DN 141:360522

TI Cardiopulmonary effects of the novel neuromuscular blocking drug GW280430A (AV430A) in dogs

AU Heerd, Paul M.; Kang, Richard; The', Andrew; Hashim, Mir; Mook, Robert J.; Savarese, John J.

CS Departments of Anesthesiology and Pharmacology, Weill Medical College of Cornell University, New York, NY, 10021, USA

SO Anesthesiology (2004), 100(4), 846-851

CODEN: ANESAV; ISSN: 0003-3022

PB Lippincott Williams & Wilkins

DT Journal

LA English

AB Background: This investigation determined the cardiopulmonary side effects of a novel nondepolarizing neuromuscular blocking drug with an ultrashort duration of action in anesthetized male beagles. Methods: The ED95 for GW280430A was first determined in four animals. These data were then used to guide bolus dosing in multiples of ED95 in six dogs instrumented for hemodynamic measurements as well as inspiratory pressure and pulmonary compliance. Cardiopulmonary data were compared before and after the conclusion of a 60- to 90-min GW280430A infusion and in response to subsequent incremental bolus dosing starting with 3.125 + ED95. An adverse response was regarded as an alteration of 10% or greater in any variable. Arterial blood was obtained for histamine anal. before and 1 min after each dose. Results: The ED95 of GW280430A was 0.064 ± 0.01 mg/kg, and stable neuromuscular blockade was maintained with infusion of 0.012 ± 0.002 mg · kg⁻¹ · min⁻¹. With the exception of a late 14% increase in heart rate, there were no cardiopulmonary changes during infusion. Bolus dosing produced no cardiopulmonary change until a decrease in mean arterial pressure was elicited in four of six dogs at 25 + ED95. This response was modest, transient, and associated with a concomitant increase in plasma histamine concentration. There were no accompanying changes indicative of direct myocardial depression, pulmonary vasoconstriction, or bronchospasm. Conclusions: These data indicate

that GW280430 does not produce demonstrable cardiovascular effects in the anesthetized dog until doses far in excess of the ED95 are administered as a bolus.

IT 213998-46-0, AV 4430A

RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(AV 4430A, AV430A; GW280430A does not produce demonstrable cardiovascular effects in anesthetized dog until dose far greater than ED95 administered as rapid i.v. bolus)

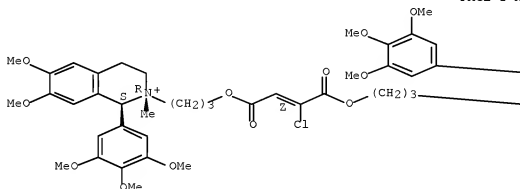
RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

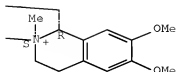
Double bond geometry as shown.

PAGE 1-A



● 2 Cl⁻

PAGE 1-B



OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)
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 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 4 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2004:362286 CAPLUS Full-text
 DN 141:360521

TI Preclinical pharmacology of GW280430A (AV430A) in the rhesus monkey and in the cat. A comparison with mivacurium

AU Savarese, John J.; Belmont, Matthew R.; Hashim, Mir A.; Mook, Robert A.; Boros, Eric E.; Samano, Vicente; Patel, Sanjay S.; Feldman, Paul L.; Schultz, Jan-Ake I.; McNulty, Michael; Spitzer, Timothy; Cohn, Douglas L.; Morgan, Philip; Wastila, William B.

CS Department of Anesthesiology, Weill Medical College of Cornell University and New York-Presbyterian Hospital, New York, NY, 10021, USA

SO Anesthesiology (2004), 100(4), 835-845
 CODEN: ANESAV; ISSN: 0003-3022

PB Lippincott Williams & Wilkins
 DT Journal
 LA English

AB Background: No replacement for succinylcholine is yet available. GW280430A (AV430A) is a representative of a new class of nondepolarizing neuromuscular blocking drugs called asym. mixed-onium chlorofumarates. It undergoes rapid degradation in plasma by chemical hydrolysis and inactivation by cysteine adduction, resulting in a very short duration of effect. The neuromuscular, cardiovascular, and autonomic pharmacol. of GW280430A is compared herein with that of mivacurium. Methods: Adult male rhesus monkeys and adult male cats were anesthetized with nitrous oxide-oxygen-halothane and chloralose-pentobarbital, resp. The neuromuscular blocking properties of GW280430A and mivacurium were compared at a stimulation rate of 0.15 Hz in the extensor digitorum of the foot (monkey) and the tibialis anterior (cat). Sympathetic responses were assayed in the cat in the nictitating membrane preparation, and vagal effects were evaluated in the cat via observation of bradycardic responses after stimulation of the cervical right vagus nerve. Results: GW280430A and mivacurium were equipotent in the monkey (ED95 was 0.06 mg/kg in each case). GW280430A was half as potent as mivacurium in the cat. The total duration of action of GW280430A was less than half that of mivacurium in the monkey; recovery slopes were more than twice as rapid. The 25-75% recovery index of GW280430A did not vary significantly after various bolus doses or infusions, averaging 1.4-1.8 min in the monkey, significantly shorter than the same time interval (4.8-5.7 min) for mivacurium. Dose ratios for autonomic vs. neuromuscular blocking properties in the cat were greater than 25 for both GW280430A and mivacurium. The ratio ED Hist:ED95 Neuromuscular Block in the monkey was significantly greater (approx. 53 vs. 13) for GW280430A, indicating approx. four times less relative prominence of the side effects of skin flushing and decrease of blood pressure, which are associated with release of histamine. Conclusions: These expts. show a much shorter neuromuscular blocking effect and much-reduced side effects in the case of GW280430A vis-a-vis mivacurium. These results, together with the novel chemical degradation of GW280430A, suggest further evaluation in human subjects.

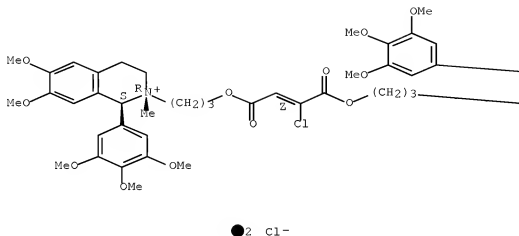
IT 213998-46-0, GW280430A
 RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (AV430A; GW280430A showed much shorter neuromuscular blocking effect and reduced side effects in anesthetized rhesus monkey, cat and chemical degradation, cardiovascular response compared with mivacurium suggest further evaluation in human subject)

RN 213998-46-0 CAPLUS

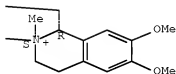
CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinol]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A



PAGE 1-B



OSC.G 7 THERE ARE 7 CAPLUS RECORDS THAT CITE THIS RECORD (7 CITINGS)
RE.CNT 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 5 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2004:362280 CAPLUS [Full-text](#)

DN 141:343098

TI Clinical pharmacology of GW280430A in humans

AU Belmont, Matthew R.; Lien, Cynthia A.; Tjan, Joseph; Bradley, Eleanor; Stein, Brenna; Patel, Sanjay S.; Savarese, John J.

CS Weill Medical College of Cornell University and New York Presbyterian Hospital, New York, NY, 10021, USA

SO Anesthesiology (2004), 100(4), 768-773

CODEN: ANESAV; ISSN: 0003-3022

PB Lippincott Williams & Wilkins

DT Journal

LA English

AB Background: An ultrashort-acting nondepolarizing neuromuscular blocking agent that could be an alternative to succinylcholine has been the focus of a concerted effort in the field of muscle relaxants. GW280430A showed a

promising pharmacodynamic profile in preclin. work and a wide margin of safety and so was selected for study in humans. Methods: Thirty-one volunteers participated in this study, which determined the dose producing 95% block (ED95) and the safety and pharmacodynamics of increasing ED95 multiples. Anesthesia was induced and maintained with propofol, midazolam, and fentanyl. Neuromuscular transmission was measured at the adductor pollicis using ulnar nerve stimulation, and responses were recorded continuously by standard mechanomyog. monitoring. Results: The ED95 for GW280430A is 0.19 mg/kg. The time to onset of 90% block ranged from 1.3 to 2.1 min, depending on the dose. Clin. durations ranged from 4.7 to 10.1 min and increased with increasing dose. Five to 95% and 25-75% recovery rates were approx. 7 and 3 min, resp., and were independent of the dose administered. Transient cardiovascular side effects were observed at doses beginning at 3 x ED95 and above and were suggestive of histamine release. Most volunteers receiving 4 x ED95 exhibited plasma histamine concns. indicative of significant histamine release. Conclusions: GW280430A has a rapid onset and ultrashort duration of action. The recovery rate is rapid, predictable, and independent of dose. Doses at least up to 2.5 x ED95 seem to be free of side effects and seem to be able to provide relaxation within 60-90 s.

IT 213998-46-0, GW280430A

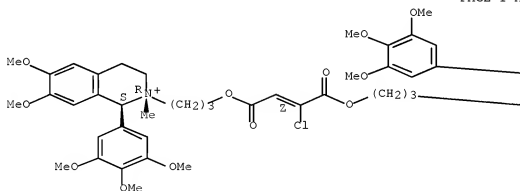
RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(GW280430A had rapid onset and ultrashort duration of action, recovery rate was rapid, predictable and independent of dose and doses at least up to 2.5 x ED95 seemed to be free of side effect and provided relaxation within 60-90 s in human)

RN 213998-46-0 CAPLUS

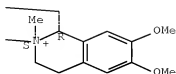
CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A

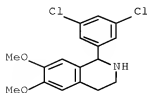


● 2 Cl⁻



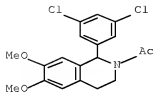
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 RE.CNT 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 6 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2004:78855 CAPLUS Full-text
 DN 140:350017
 TI Synthesis and anticonvulsant properties of tetrahydroisoquinoline derivatives
 AU Gitto, Rosaria; Caruso, Roberta; Orlando, Valerie; Quartarone, Silvana; Barreca, Maria Letizia; Ferreri, Guido; Russo, Emilio; De Sarro, Giovambattista; Chimirri, Alba
 CS Dipartimento Farmaco-Chimico, Universita di Messina, Messina, 98168, Italy
 SO Farmaco (2004), 59(1), 7-12
 CODEN: FRMCE8; ISSN: 0014-827X
 PB Editions Scientifiques et Medicales Elsevier
 DT Journal
 LA English
 OS CASREACT 140:350017
 AB As a follow up of our previous structure-activity relationship and mol. modeling studies, we synthesized a novel series of 1-aryl-6,7-dimethoxy-1,2,3,4-tetrahydroisoquinoline derivs. as potential non-competitive AMPA receptor antagonists. When tested for their ability to prevent sound-induced seizures in DBA/2 mice, some of these novel compds. showed high anticonvulsant potency.
 IT 682763-23-1P 682763-30-0P
 RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (synthesis and anticonvulsant properties of tetrahydroisoquinoline derivs. that act as non-competitive AMPA receptor antagonists)
 RN 682763-23-1 CAPLUS
 CN Isoquinoline, 1-(3,5-dichlorophenyl)-1,2,3,4-tetrahydro-6,7-dimethoxy- (CA INDEX NAME)



RN 682763-30-0 CAPLUS

CN Ethanone, 1-[1-(3,5-dichlorophenyl)-3,4-dihydro-6,7-dimethoxy-2(1H)-isoquinolinyl]- (CA INDEX NAME)



OSC.G 13 THERE ARE 13 CAPLUS RECORDS THAT CITE THIS RECORD (13 CITINGS)
RE.CNT 31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 7 OF 61 CAPLUS COPYRIGHT 2009 ACS ON STN

AN 2003:347314 CAPLUS Full-text

DN 139:78440

TI Neuromuscular Blocking Activity and Therapeutic Potential of Mixed-Tetrahydroisoquinolinium Halofumarates and Halosuccinates in Rhesus Monkeys

AU Boros, Eric E.; Samano, Vicente; Ray, John A.; Thompson, James B.; Jung, David K.; Kaldor, Istvan; Koble, Cecilia S.; Martin, Michael T.; Styles, Virgil L.; Mook, Robert A., Jr.; Feldman, Paul L.; Savarese, John J.; Belmont, Matthew R.; Bigham, Eric C.; Boswell, G. Evan; Hashim, Mir A.; Patel, Sanjay S.; Wisowaty, James C.; Bowers, Gary D.; Moseley, Caroline L.; Walsh, John S.; Reese, Mindy J.; Rutkowske, Randy D.; Seffler, Andrea M.; Spitzer, Timothy D.

CS GlaxoSmithKline Research & Development, Research Triangle Park, NC, 27709, USA

SO Journal of Medicinal Chemistry (2003), 46(12), 2502-2515

CODEN: JMCMAR; ISSN: 0022-2623

PB American Chemical Society

DT Journal

LA English

OS CASREACT 139:78440

AB Structure-activity relationships in rhesus monkeys for a novel mixed-onium class of ultra-short-acting nondepolarizing tetrahydroisoquinolinium neuromuscular blockers (NMBs) are described. Bis-onium chlorofumarate 20a with (1R,2S)-benzyltetrahydroisoquinolinium groups was a potent lead compound (ED95 = 0.079 mg/kg) with an ultra-short duration of NMB effect (7.1 min) and a selectivity index (SI: defined as a ratio of the cardiovascular threshold dose to the ED95) similar to that of mivacurium. The mean threshold dose for cardiovascular effects with 20a was .apprx.20 times its ED95 value (SI = 20). A novel mixed-onium analog of was prepared by replacing the benzyltetrahydroisoquinolinium group distal to the fumarate chlorine atom with a (1S,2R)-phenyltetrahydroisoquinolinium moiety. The resulting mixed-onium chlorofumarate 24a displayed good NMB potency (ED95 = 0.063 mg/kg), ultra-short duration of action (5.6 min) and an improved selectivity index (SI = 57). Several other mixed-onium derivs. containing octanedioate (ED95 = 0.103 mg/kg), difluorosuccinate (ED95 = 0.056 mg/kg), and fluorofumarate (ED95 = 0.137 mg/kg) linkers were also potent, ultra-short-acting NMBs with good to excellent selectivity index values (SI = 37 - 96). Octanedioate was longer acting at higher doses compared to difluorosuccinate and chlorofumarate. Durations of NMB effect following a 0.4 mg/kg bolus dose (100% block) of octanedioate, difluorosuccinate, fluorofumarate and were 16.9, 13.0, and 10.0 min, resp. Recovery time for mixed-onium chlorofumarate following a 1 h

continuous infusion at 10 - 20 µg/kg/min (95 - 100% block) was .apprx.5 min which is similar to that observed following a 0.2 mg/kg bolus dose of this compound and indicates a lack of cumulative effects. Preliminary studies with chlorofumarate in whole human blood revealed that mixed-onium thiazolidine was the major metabolite and that plasma cholinesterases do not play the primary role in duration of NMB effect. The NMB properties of chlorofumarate in rhesus monkeys led to its clin. evaluation as a possible alternative to succinylcholine.

| | | | |
|----|--------------|--------------|--------------|
| IT | 213998-54-0P | 213998-59-5P | 213998-64-2P |
| | 213998-65-3P | 213998-67-5P | 213998-69-7P |
| | 213998-71-1P | 213998-31-3P | 213998-82-4P |
| | 213998-83-5P | 213998-84-6P | 213998-26-3P |
| | 213999-39-4P | 213999-50-9P | 213999-51-0P |
| | 213999-52-1P | 347364-95-6P | 552319-33-2P |
| | 552319-34-3P | 552319-36-5P | |

RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(neuromuscular blocking activity and therapeutic potential of mixed-tetrahydroisoquinolinium halofumarates and halosuccinates in rhesus monkeys)

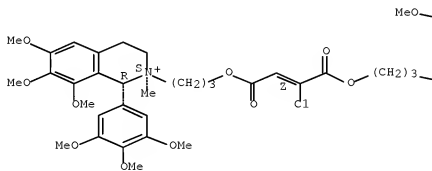
RN 213998-54-0 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

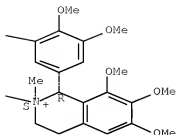
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A



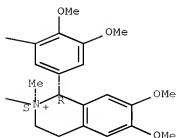
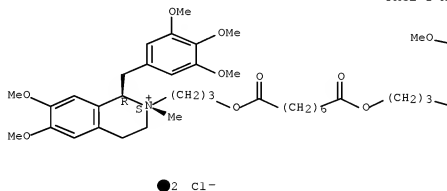
● 2 Cl⁻



RN 213998-59-5 CAPLUS

CN Isoquinolinium, 2-[3-[[1,8-dioxo-8-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]octyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

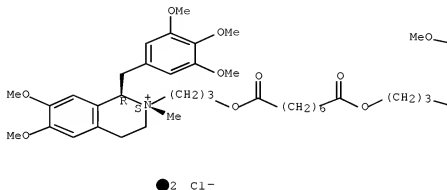


RN 213998-64-2 CAPLUS

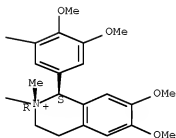
CN Isoquinolinium, 2-[3-[[1,8-dioxo-8-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]octyl]oxy]propyl]-1,2,3,4-

tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-,
dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

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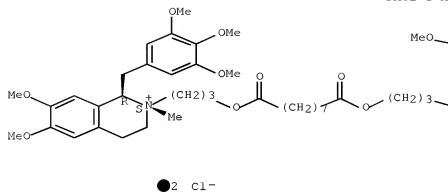
PAGE 1-B



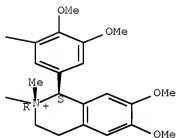
| | |
|----|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| CN | <p> Isoquinolinium 2-[3-[[1,9-dioxo-9-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]nonyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME) </p> |
|----|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|

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PAGE 1-B

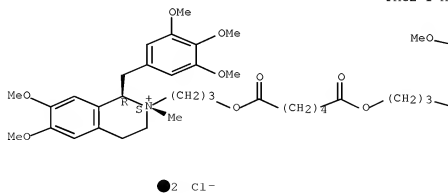


RN 213998-67-5 CAPLUS

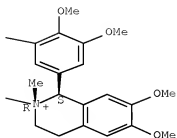
CN Isoquinolinium, 2-[3-[[1,6-dioxo-6-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]hexyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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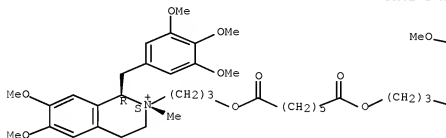


RN 213998-69-7 CAPLUS

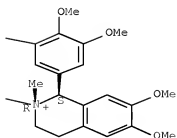
CN Isoquinolinium, 2-[3-[[1,7-dioxo-7-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]heptyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

● 2 Cl⁻

PAGE 1-B



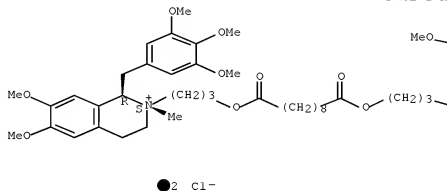
RN 213998-71-1 CAPLUS

CN Isoquinolinium, 2-[3-[[1,10-dioxo-10-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-

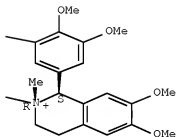
dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]decyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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PAGE 1-B



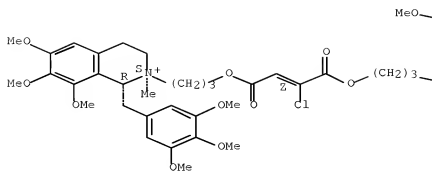
RN 213998-81-3 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

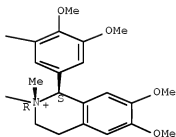
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

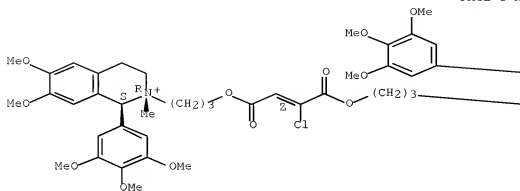
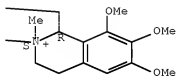
PAGE 1-B



RN 213998-82-4 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

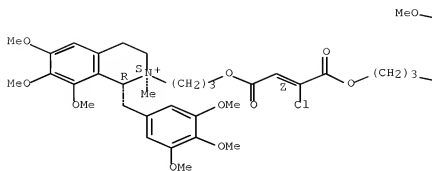
RN 213998-83-5 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxypropyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)]

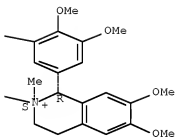
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

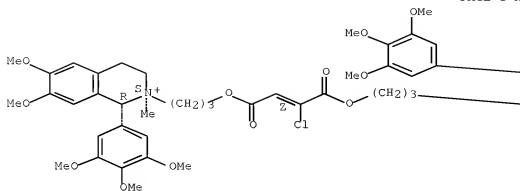
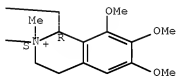
PAGE 1-B



RN 213998-84-6 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)-
(CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

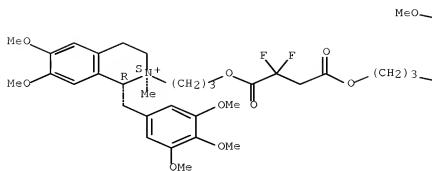
● 2 Cl⁻

RN 213999-26-9 CAPLUS

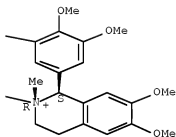
CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-4-[3-[(1S)-3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-2(1H)-isoquinolinyl]propoxy]-1,4-dioxo-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride, hydrochloride (1:1:1), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

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● 2 Cl⁻

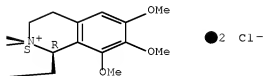
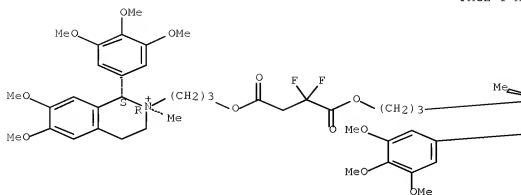
PAGE 1-B



RN 213999-50-9 CAPLUS

CN Isoquinolinium, 2-[3-[2,2-difluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

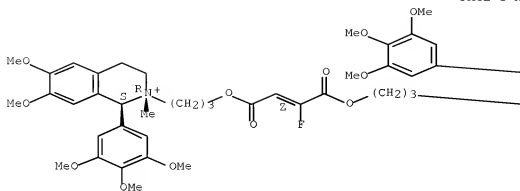
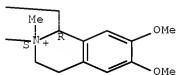


RN 213999-51-0 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

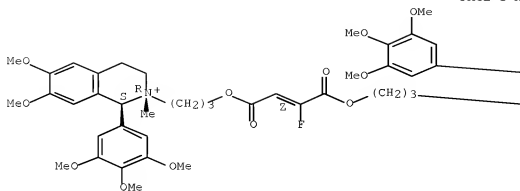
● 2 Cl⁻

RN 213999-52-1 CAPLUS

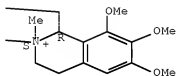
CN Isoquinolinium, 2-[3-[[(2Z)-2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]propoxy]-2-buten-1-yl]oxypropyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)]

Absolute stereochemistry.

Double bond geometry as shown.



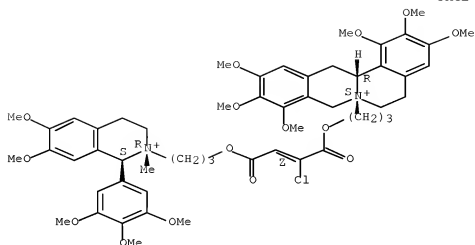
● 2 Cl⁻



RN 347384-95-6 CAPLUS

CN 6H-Dibenzo[a,g]quinolizinium, 7-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-5,8,13,13a-tetrahydro-1,2,3,9,10,11-hexamethoxy-, dichloride, (7S,13aR)-(9CI) (CA INDEX NAME)

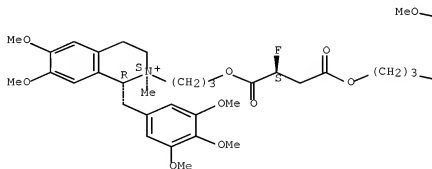
Absolute stereochemistry.
Double bond geometry as shown.



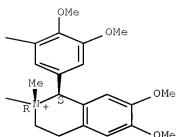
RN 552319-33-2 CAPLUS

CN Isoquinolinium, 2-[3-[(2S)-2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



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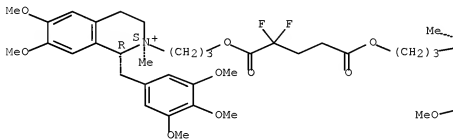


RN 552319-34-3 CAPLUS

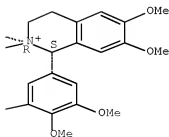
CN Isoquinolinium, 2-[3-[[2,2-difluoro-1,5-dioxo-5-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]pentyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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● 2 Cl⁻

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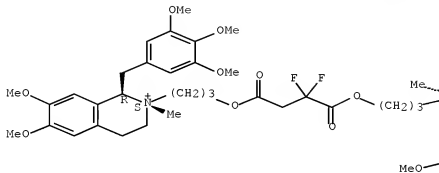
RN 552319-36-5 CAPLUS

CN Isoquinolinium, 2-[3-[3,3-difluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-

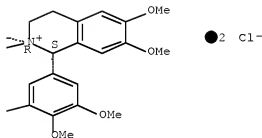
trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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PAGE 1-B



IT 213998-45-9P 213998-46-0P 213998-47-1P
213998-48-2P 213998-53-9P 213998-57-3P
213998-58-4P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(neuromuscular blocking activity and therapeutic potential of mixed-tetrahydroisoquinolinium halofumarates and halosuccinates in rhesus monkeys)

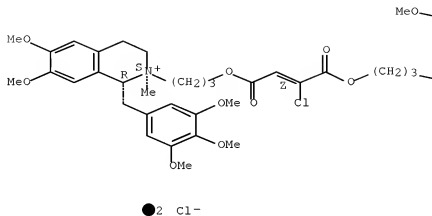
RN 213998-45-9 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

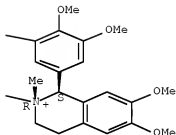
Absolute stereochemistry.

Double bond geometry as shown.

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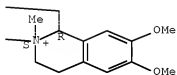
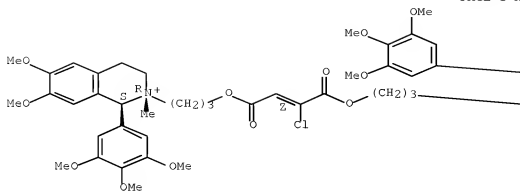


RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

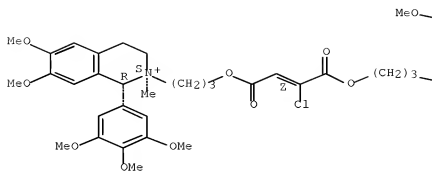


RN 213998-47-1 CAPLUS

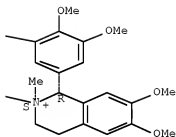
CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.



● 2 Cl-

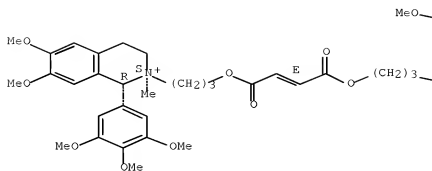
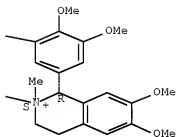


RN 213998-48-2 CAPLUS

CN Isoquinolinium, 2,2'-[[(2E)-1,4-dioxo-2-butene-1,4-diyl]bis (oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

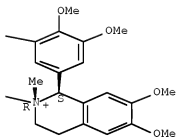
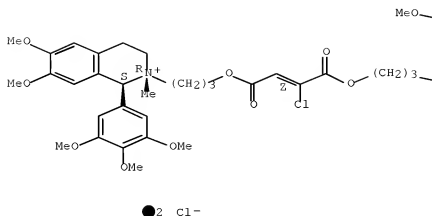
● 2 Cl⁻

RN 213998-53-9 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,1'S,2R,2'R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

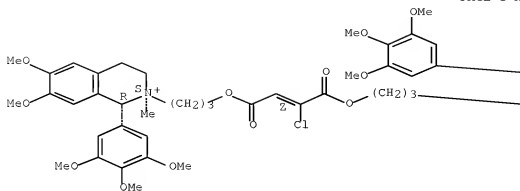
Double bond geometry as shown.



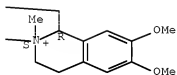
RN 213998-57-3 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.



● 2 Cl⁻



IT 198401-05-7P 220408-26-4P 552319-26-3P

552319-28-5P 552319-32-1P

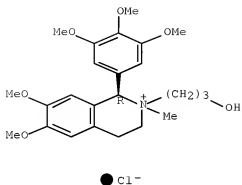
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(neuromuscular blocking activity and therapeutic potential of mixed-tetrahydroisoquinolinium halofumarates and halosuccinates in rhesus monkeys)

RN 198401-05-7 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1R)- (CA INDEX NAME)

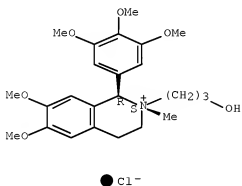
Absolute stereochemistry.



RN 220408-26-4 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1R,2S)- (CA INDEX NAME)

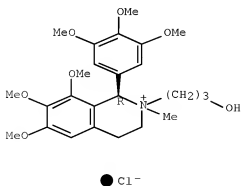
Absolute stereochemistry.



RN 552319-26-3 CAPLUS

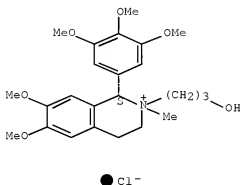
CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7,8-trimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1R)- (CA INDEX NAME)

Absolute stereochemistry.



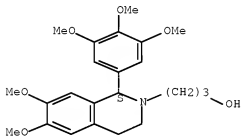
RN 552319-28-5 CAPLUS
 CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S)- (CA INDEX NAME)

Absolute stereochemistry.



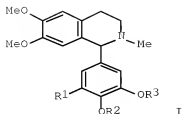
RN 552319-32-1 CAPLUS
 CN 2(1H)-Isoquinolinepropanol, 3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry.



OSC.G 9 THERE ARE 9 CAPLUS RECORDS THAT CITE THIS RECORD (10 CITINGS)
 RE.CNT 50 THERE ARE 50 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 8 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2003:281089 CAPLUS [Full-text](#)
 DN 139:180218
 TI A one-pot synthesis of (±) cryptostylin
 AU Ruchirawat, Somsak; Bhavakul, Vanida; Chaisupakitsin, Malinee
 CS Chulabhorn Research Institute, Bangkok, 10210, Thailand
 SO Synthetic Communications (2003), 33(4), 621-625
 CODEN: SYNCAV; ISSN: 0039-7911
 PB Marcel Dekker, Inc.
 DT Journal
 LA English
 OS CASREACT 139:180218
 GI

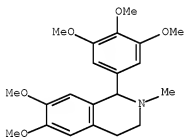


AB A one-pot synthesis of cryptostylinines I (R1 = H, OMe; R2, R3 = Me; R2-R3 = CH2) via the Pictet-Spengler reaction is reported.

IT 22324-83-0P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (one-pot preparation of (±) cryptostyline alkaloids from homoveratrylamine and aromatic aldehydes via Pictet-Spengler reaction and N-methylation with formaldehyde)

RN 22324-83-0 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)

RE.CNT 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 9 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2002:704029 CAPLUS Full-text

DN 138:231172

TI The pharmacology of GW280430A: a new nondepolarizing neuromuscular blocking agent

AU Lien, Cynthia A.

CS Department of Anesthesiology, New York Presbyterian Hospital, Weill Medical College of Cornell University, New York, NY, USA

SO Seminars in Anesthesia, Perioperative Medicine and Pain (2002), 21(2), 86-91

CODEN: SAPPFJ

PB W. B. Saunders Co.

DT Journal; General Review

LA English

AB A review. GW280430A is a new nondepolarizing neuromuscular blocking agent, which is different from any other used either currently or in the past. It is a potent nondepolarizing agent that has a rapid onset of effect comparable in early volunteer trials to that of succinylcholine. Its unique chemical structure is responsible not only for its potency and nondepolarizing

neuromuscular blocking activity but also for its means of elimination from the plasma. Although other relaxants, such as mivacurium, undergo hydrolysis by plasma esterases, no other compound undergoes chemical hydrolysis and cysteine adduction. These reactions happen extremely quickly and are likely responsible for the ultrashort duration of action of GW280430A. If the results of further trials of this compound in patients undergoing surgical procedures are similar to those of the earlier volunteer trials, GW280430A promises to be the nondepolarizing equivalent of succinylcholine. Its development may allow for greater safety with the use of nondepolarizing neuromuscular blocking agents. With its ultrashort duration of action, unacceptable levels of neuromuscular blockade after extubation of the trachea and all associated adverse sequelae, such as decreased hypoxic drive to breathe, respiratory failure, and increased incidence of aspiration, may become things of the past.

IT 213998-46-0, GW 280430A

RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); PKT (Pharmacokinetics); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmacol. of new nondepolarizing neuromuscular blocking agent GW280430A in laboratory animals and humans)

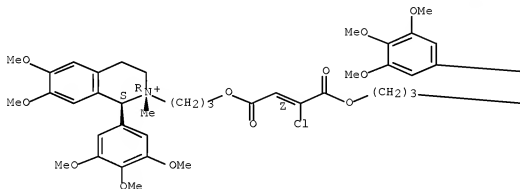
RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

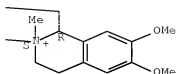
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A



● 2 Cl⁻



OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)
 RE.CNT 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 10 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2002:226386 CAPLUS [Full-text](#)

DN 137:299691

TI Preformulation studies for an ultrashort-acting neuromuscular blocking agent GW280430A. I. Buffer and cosolvent effects on the solution stability

AU Zhu, Haijian; Meserve, Kathy; Floyd, Alison

CS GlaxoSmithKline Inc., Research Triangle Park, NC, 27709, USA

SO Drug Development and Industrial Pharmacy (2002), 28(2), 135-142

CODEN: DDIPD8; ISSN: 0363-9045

PB Marcel Dekker, Inc.

DT Journal

LA English

AB GW280430A is an ultrashort-acting neuromuscular blocking agent targeted at muscle relaxation to facilitate surgical intubation. The objective of this study was to study the buffer and cosolvent effects on the solution stability of GW280430A. The buffer catalytic effect was examined in citrate, malate, tartrate, and glycine by measuring the rate of degradation of GW280430A (0.2 mg/mL) at constant pH (3), ionic strength (0.15M), and various buffer concns. (0.01-0.05M). The temperature dependence of the buffer catalytic effect and the degradation of the GW280430A in cosolvent (ethanol, propylene glycol, polyethylene glycol 400, N,N-dimethylacetamide)/water mixts. were studied at 40, 50, and 60°. The loss of parent drug was monitored by reverse-phase HPLC. The degradation of GW280430A followed first-order kinetics in all buffer solns. Significant buffer-catalyzed hydrolysis of GW280430A was observed with citrate, tartrate, and malate buffers, but not in glycine-buffered solns. The activation energies in all buffered drug solns. ranged from 70 to 80 kJ/mol and decreased with increasing buffer concentration. GW280430A degradation was primarily through ester hydrolysis and followed first-order kinetics in aqueous solns. In cosolvent/water mixts., new degradation products were observed, indicating a chemical reaction between GW280430A and cosolvents. The reaction activation energies in the cosolvent/water mixts. ranged 75-85 kJ/mol, with the longest t_{0.9} at 5° equal to approx. 12 mo and at 25° equal to 36 days. Consideration should be given to the incorporation of glycine or a low concentration of citrate, malate, or tartrate buffer in the parenteral formulation development of GW280430A. Cosolvents prolonged the predicted t_{0.9} for GW280430A in solution, but the enhancement was not significant enough to pursue a liquid formulation.

IT 213998-46-G, GW 280430A

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(buffer and cosolvent effects on solution stability of neuromuscular blocker GW280430A)

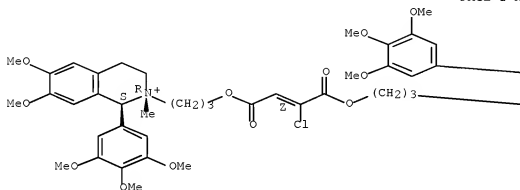
RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

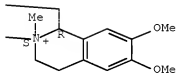
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

PAGE 1-B



OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)
 RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD
 DN 137:284090 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 11 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2002:115984 CAPLUS [Full-text](#)

DN 137:284090

TI Solid state characterization of an neuromuscular blocking agent-GW280430A

AU Zhu, Haijian; Sacchetti, Mark

CS GlaxoSmithKline Inc., Research Triangle Park, NC, 27709, USA

SO International Journal of Pharmaceutics (2002), 234(1-2), 19-23

CODEN: IJPHDE; ISSN: 0378-5173

PB Elsevier Science B.V.

DT Journal
 LA English
 AB GW280430A is an ultrashort-acting neuromuscular blocking agent and is targeted for muscle relaxation as part of the intubation surgical procedure. The objective of this study was to perform solid state characterization on GW280430A and to evaluate the relationship between water content and glass transition temperature (Tg). GW280430A was characterized by differential scanning calorimetry, thermogravimetric anal., powder x-ray diffraction (PXRD), microscopy and moisture sorption. The effect of water content on the Tg of GW280430A was evaluated by equilibrating the material over saturated salt solns. at a range of relative humidities (6.4–72.6%) and determining the Tg by DSC using hermetically sealed aluminum pans. GW280430A undergoes dehydration at 40°, glass transition at 130° and decomposition at 190° by DSC. By PXRD and moisture sorption, GW280430A is an amorphous material and deliquesces at about 70% RH at room temperature. Water acts as a potent plasticizer for GW280430A and the Tg decreases significantly as the water content increases. No measurable decomposition of GW280430A was observed after 4 wk at 40°/75% RH.

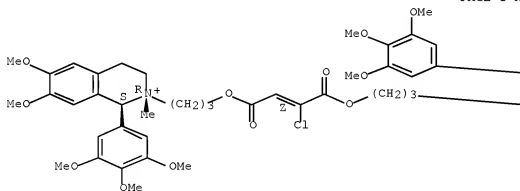
IT 213998-46-0, GW 280430A
 RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (solid state characterization of an neuromuscular blocking agent GW280430A)

RN 213998-46-0 CAPLUS

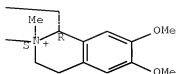
CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.

PAGE 1-A



● 2 Cl⁻



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
 RE.CNT 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 12 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2001:472522 CAPLUS [Full-text](#)

DN 135:66254

TI pharmaceutical formulations containing histamine releasers

IN Floyd, Alison G.; Hashim, Mir A.; Lin, Peiyuan; Mook, Robert A.; Sefler, Andrea

PA Glaxo Group Ltd., UK

SO PCT Int. Appl., 91 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

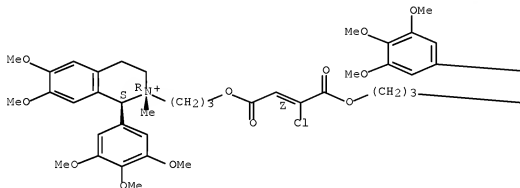
| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| PI | WO 2001045741 | A2 | 20010628 | WO 2000-US33772 | 20001213 <-- |
| | WO 2001045741 | A3 | 20011206 | | |
| | W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | |
| | RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| | CA 2394794 | A1 | 20010628 | CA 2000-2394794 | 20001213 <-- |
| | CA 2394794 | C | 20090825 | | |
| | EP 1239884 | A2 | 20020918 | EP 2000-984305 | 20001213 <-- |
| | EP 1239884 | B1 | 20070411 | | |
| | R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | |
| | JP 2003518074 | T | 20030603 | JP 2001-546680 | 20001213 <-- |
| | AU 779889 | B2 | 20050217 | AU 2001-20945 | 20001213 |
| | AT 359092 | T | 20070515 | AT 2000-984305 | 20001213 |
| | ES 2284548 | T3 | 20071116 | ES 2000-984305 | 20001213 |
| | US 20030096839 | A1 | 20030522 | US 2002-149722 | 20020613 <-- |
| | US 6911455 | B2 | 20050628 | | |
| | MX 2002006266 | A | 20021205 | MX 2002-6266 | 20020621 <-- |
| PRAI | US 1999-171696P | P | 19991222 | | |
| | WO 2000-US33772 | W | 20001213 | | |

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

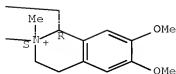
- AB The invention relates to pharmaceutical formulations and methods for preparing pharmaceutical formulations of histamine releasers. The present invention provides methods for determining the concentration of physiologically acceptable excipient for use in the formulations of invention. Methods for suppressing pharmaceutically-induced histamine release by administering the formulations to an animal are also provided. A kit useful for preparing pharmaceutical formulations of histamine releasers is also described. The percent inhibition of histamine release by formulations containing a tetrahydroisoquinoline derivative and citric acid, an excipient (12.5 mg/mL), at pH 3 was 90.8.
- IT 213998-46-0
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (pharmaceutical formulations containing histamine releasers)
- RN 213998-46-0 CAPLUS
- CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

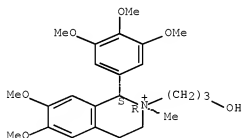
PAGE 1-B



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
 RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 13 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2001:290361 CAPLUS Full-text
 DN 135:77006
 TI Stereocontrolled Synthesis of cis-Dibenzoquinolizinium Chlorofumarates:
 Curare-Like Agents of Ultrashort Duration
 AU Kaldor, Istvan; Feldman, Paul L.; Mook, Robert A., Jr.; Ray, John A.;
 Samano, Vicente; Sefler, Andrea M.; Thompson, James B.; Travis, Benjamin
 R.; Boros, Eric E.
 CS Division of Medicinal Chemistry, GlaxoSmithKline Research & Development,
 Research Triangle Park, NC, 27709, USA
 SO Journal of Organic Chemistry (2001), 66(10), 3495-3501
 CODEN: JOCEAH; ISSN: 0022-3263
 PB American Chemical Society
 DT Journal
 LA English
 OS CASREACT 135:77006
 AB Cis-Dibenzoquinoliziniumpropanols were prepared stereoselectively and were
 transformation into bis- and mixed-onium chlorofumarates. The title comps.
 displayed curare-like effects of ultrashort duration in rhesus monkeys.
 IT 213999-53-2
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (stereocontrolled synthesis of cis-dibenzoquinolizinium chlorofumarates,
 curare-like agents of ultrashort duration)
 RN 213999-53-2 CAPLUS
 CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-
 methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2R)- (CA INDEX
 NAME)

Absolute stereochemistry.



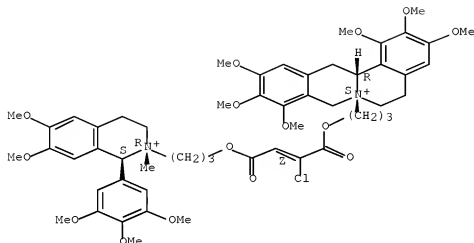
● Cl⁻

IT 347384-95-6P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (stereocontrolled synthesis of cis-dibenzoquinolizinium chlorofumarates,
 curare-like agents of ultrashort duration)
 RN 347384-95-6 CAPLUS
 CN 6H-Dibenzo[a,g]quinolizinium, 7-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-
 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-
 trimethoxyphenyl)methyl]isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-
 5,8,13,13a-tetrahydro-1,2,3,9,10,11-hexamethoxy-, dichloride, (7S,13aR)-

(9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A



PAGE 2-A

● 2 Cl-

OSC.G 14 THERE ARE 14 CAPLUS RECORDS THAT CITE THIS RECORD (14 CITINGS)
RE.CNT 29 THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 14 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
AN 2000:553442 CAPLUS [Full-text](#)
DN 133:168383
TI Pharmaceutical compositions containing nicotine or a ligand of nicotine
receptors and a monamine oxidase inhibitor and their use for treating
tobacco withdrawal symptoms
IN Caille, Dominique; George, Pascal; Jegham, Samir; Robineau, Pascale;
Scatton, Bernard; Zivkovic, Branimir
PA Sanofi-Synthelabo, Fr.
SO PCT Int. Appl., 37 pp.
CODEN: PIXXD2
DT Patent
LA French
FAN.CNT 1

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| WO 2000045846 | A1 | 20000810 | WO 2000-FR193 | 20000128 <-- |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |

RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

| | | | | |
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| FR 2788982 | B1 | 20020802 | | |
| CA 2361437 | A1 | 20000810 | CA 2000-2361437 | 20000128 <-- |
| EP 1150715 | A1 | 20011107 | EP 2000-901660 | 20000128 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| HU 2002001279 | A2 | 20020928 | HU 2002-1279 | 20000128 <-- |
| JP 2002536342 | T | 20021029 | JP 2000-596965 | 20000128 <-- |
| MX 2001007812 | A | 20020108 | MX 2001-7812 | 20010802 <-- |
| PRAI FR 1999-1144 | A | 19990202 | | |
| WO 2000-FR193 | W | 20000128 | | |

OS MARPAT 133:168383

AB The invention concerns novel pharmaceutical compns. containing nicotine or a ligand of nicotine receptors and a monamine oxidase inhibitor designed for treating tobacco withdrawal symptoms. A bilayer tablet contained bexloxadone 5, lactose 66, microcryst. cellulose 20, povidone 4, crospovidone 4, and magnesium stearate 1% in the first layer, and nicotine polarcricilex 5, microcryst. cellulose 20 povidone 4, hydroxypropyl Me cellulose 25, magnesium stearate 1, and lactose q.s. 100% in the second layer.

IT 213998-46-0, GW 280430

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compns. containing nicotine or ligand of nicotine receptors and monamine oxidase inhibitor and their use for treating tobacco withdrawal symptoms)

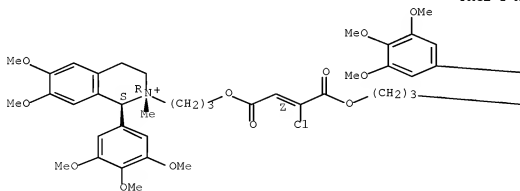
RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

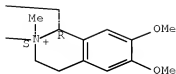
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A



● 2 Cl-



OSC.G 9 THERE ARE 9 CAPLUS RECORDS THAT CITE THIS RECORD (9 CITINGS)
 RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 15 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1999:757502 CAPLUS Full-text

DN 132:107858

TI Synthesis of Ultra-Short-Acting Neuromuscular Blocker GW 0430: A
 Remarkably Stereo- and Regioselective Synthesis of Mixed
 Tetrahydroisoquinolinium Chlorofumarates

AU Samano, Vicente; Ray, John A.; Thompson, James B.; Mook, Robert A., Jr.;
 Jung, David K.; Koble, Cecilia S.; Martin, Michael T.; Bigham, Eric C.;
 Regitz, Craig S.; Feldman, Paul L.; Boros, Eric E.

CS Department of Chemistry, Glaxo Wellcome Research and Development, Research
 Triangle Park, NC, 27709, USA

SO Organic Letters (1999), 1(12), 1993-1996

CODEN: ORLEF7; ISSN: 1523-7060

PB American Chemical Society

DT Journal

LA English

OS CASREACT 132:107858

AB The stereo- and regioselective synthesis of ultra-short-acting nondepolarizing
 neuromuscular blocker GW 0430 is described. Key steps involved the
 enantioselective transfer hydrogenation of an imine employing Noyori's
 catalyst, stereoselective crystallization and methanolysis of trans-betaines,
 and stereo- and regioselective trans elimination of hydrogen chloride. The
 latter transformation allowed complete control of the position of the chloro
 substituent and stereochem. at the double bond of the linker.

IT 213998-45-9P 213998-46-0P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); SPN (Synthetic preparation); BIOL (Biological
 study); PREP (Preparation)

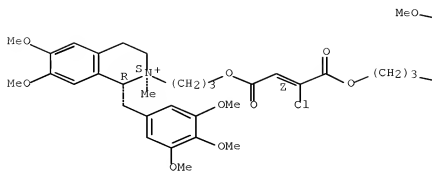
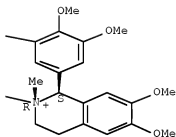
(stereo- and regioselective preparation of mixed tetrahydroisoquinolinium
 chlorofumarates)

RN 213998-45-9 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-
 tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-
 trimethoxyphenyl)isoquinolinyl]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-
 tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-,
 dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

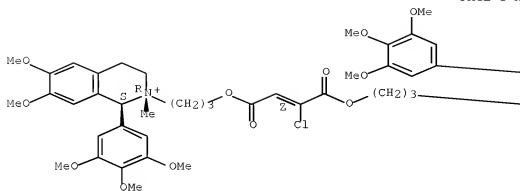
Double bond geometry as shown.

● 2 Cl⁻

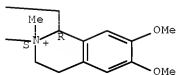
RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.



● 2 Cl-



IT 22325-16-2P 213999-53-2P 213999-54-3P

214191-49-8P 255821-84-2P

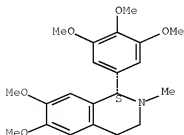
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(stereo- and regioselective preparation of mixed tetrahydroisoquinolinium chlorofumarates)

RN 22325-16-2 CAPLUS

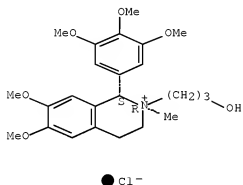
CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



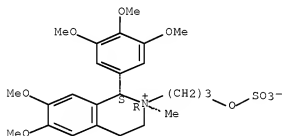
RN 213999-53-2 CAPLUS
 CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2R)- (CA INDEX NAME)

Absolute stereochemistry.



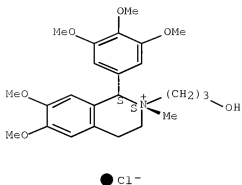
RN 213999-54-3 CAPLUS
 CN Isoquinolinium, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-2-[3-(sulfooxy)propyl]-1-(3,4,5-trimethoxyphenyl)-, inner salt, (1S,2R)- (CA INDEX NAME)

Absolute stereochemistry.



RN 214191-49-8 CAPLUS
 CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2S)- (CA INDEX NAME)

Absolute stereochemistry.



RN 255821-84-2 CAPLUS

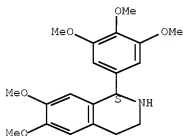
CN Formic acid, compd. with (1S)-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)isoquinoline (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 32886-69-4

CMF C20 H25 N O5

Absolute stereochemistry.



CM 2

CRN 64-18-6

CMF C H2 O2



OSC.G 15 THERE ARE 15 CAPLUS RECORDS THAT CITE THIS RECORD (15 CITINGS)

RE.CNT 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 16 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1999:173738 CAPLUS [Full-text](#)

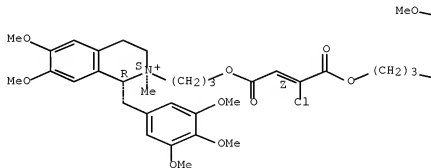
DN 130:281969

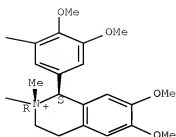
TI Bis- and Mixed-Tetrahydroisoquinolinium Chlorofumarates: New Ultra-Short-Acting Nondepolarizing Neuromuscular Blockers. [Erratum to

- document cited in CA130:168224]
- AU Boros, Eric E.; Bigham, Eric C.; Boswell, G. Evan; Mook, Robert A., Jr.; Patel, Sanjay S.; Savarese, John J.; Ray, John A.; Thompson, James B.; Hashim, Mir A.; Wisowaty, James C.; Feldman, Paul L.; Samano, Vicente
- CS Glaxo Wellcome Research and Development, Research Triangle Park, NC, 27709, USA
- SO Journal of Medicinal Chemistry (1999), 42(6), 1114
CODEN: JMCMAR; ISSN: 0022-2623
- PB American Chemical Society
- DT Journal
- LA English
- AB On page 207, under Neuromuscular Pharmacol., sentence 1, line 3, "or" should be replaced with "and": Rhesus monkeys (adult males, 8-15 kg) were anesthetized with ketamine (5 mg/kg, i.m.) and sodium pentobarbital (205 mg/kg, iv).
- IT 213998-45-9P 213998-46-0P 213998-47-1P
213998-48-2P 213998-53-9P 213998-57-3P
213998-58-4P
- RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)
- (bis- and mixed-tetrahydroisoquinolinium chlorofumarates as ultra-short-acting nondepolarizing neuromuscular blockers (Erratum))
- RN 213998-45-9 CAPLUS
- CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

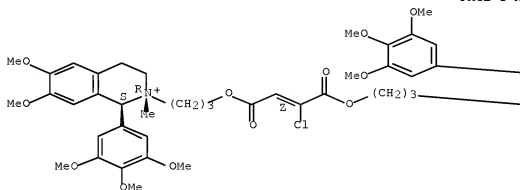
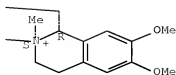


RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

● 2 Cl⁻

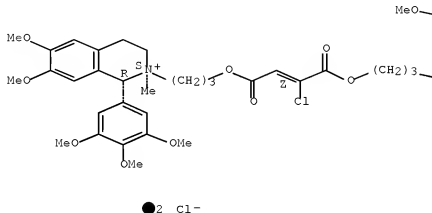
RN 213998-47-1 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

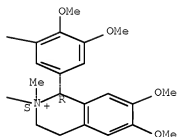
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A



PAGE 1-B

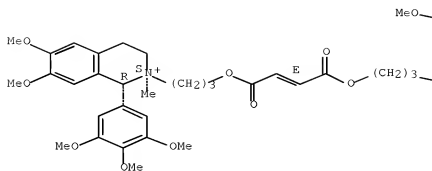
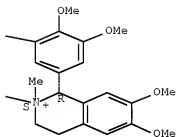


RN 213998-48-2 CAPLUS

CN Isoquinolinium, 2,2'-[[(2E)-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

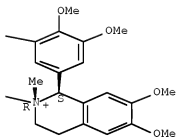
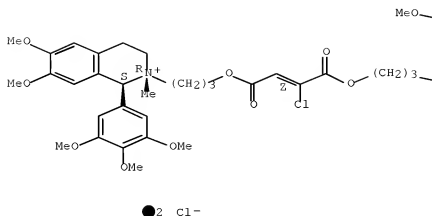
● 2 Cl⁻

RN 213998-53-9 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,1'S,2R,2'R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

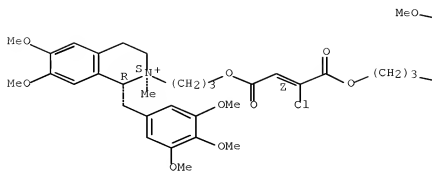
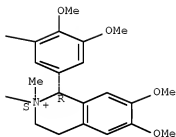
Double bond geometry as shown.



RN 213998-57-3 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

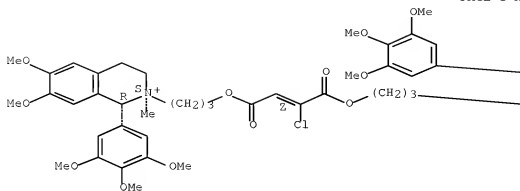
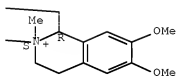
Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

RN 213998-58-4 CAPLUS

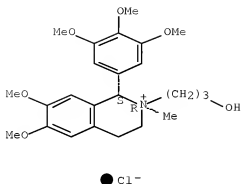
CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

IT 213999-53-2 220408-26-4
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (bis- and mixed-tetrahydroisoquinolinium chlorofumarates as
 ultra-short-acting nondepolarizing neuromuscular blockers (Erratum))
 RN 213999-53-2 CAPLUS
 CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-
 methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2R)- (CA INDEX
 NAME)

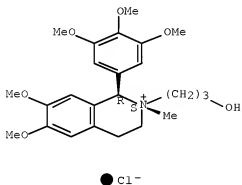
Absolute stereochemistry.



RN 220408-26-4 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.



OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)

L5 ANSWER 17 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1999:8655 CAPLUS [Full-text](#)

DN 130:168224

TI Bis- and Mixed-Tetrahydroisoquinolinium Chlorofumarates: New Ultra-Short-Acting Nondepolarizing Neuromuscular Blockers

AU Boros, Eric E.; Bigham, Eric C.; Boswell, G. Evan; Mook, Robert A., Jr.; Patel, Sanjay S.; Savarese, John J.; Ray, John A.; Thompson, James B.; Hashim, Mir A.; Wisowaty, James C.; Feldman, Paul L.; Samano, Vicente

CS Glaxo Wellcome Research and Development, Research Triangle Park, NC, 27709, USA

SO Journal of Medicinal Chemistry (1999), 42(2), 206-209
CODEN: JMCMAR; ISSN: 0022-2623

PB American Chemical Society

DT Journal

LA English

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Title compds. such as I (n = 0, 1; X = H, Cl) were prepared by reaction of the isoquinolinium headgroups with chlorofumaryl or fumaryl chloride in dichloroethane. Potency values (ED95), onset times, and duration of neuromuscular blocking action were measured in rhesus monkeys.

IT 213998-45-9P 213998-46-0P 213998-47-1P
213998-48-2P 213998-53-9P 213998-57-3P
213998-58-4P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

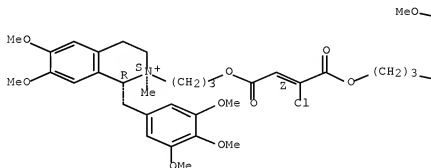
(bis- and mixed-tetrahydroisoquinolinium chlorofumarates as ultra-short-acting nondepolarizing neuromuscular blockers)

RN 213998-45-9 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (SCI) (CA INDEX NAME)

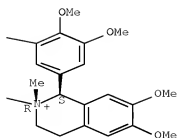
Absolute stereochemistry.
Double bond geometry as shown.

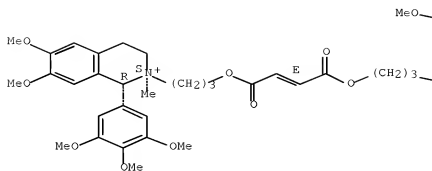
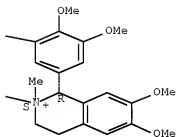
PAGE 1-A



● 2 Cl⁻

PAGE 1-B



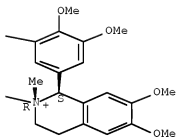
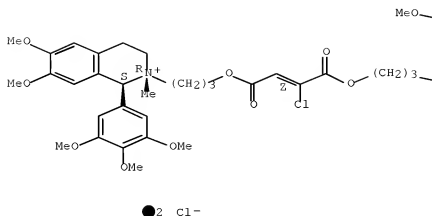
● 2 Cl⁻

RN 213998-53-9 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,1'S,2R,2'R)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

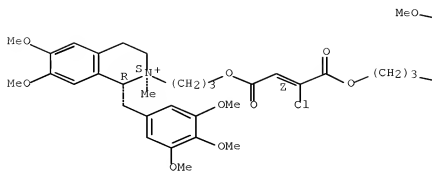
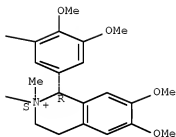
Double bond geometry as shown.



RN 213998-57-3 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)-(9CI) (CA INDEX NAME)

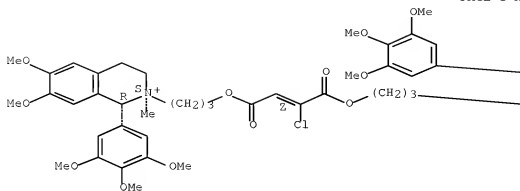
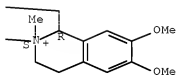
Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

RN 213998-58-4 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

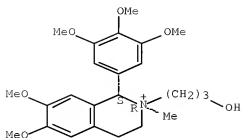
IT 213999-53-2 220408-26-4

RL: RCT (Reactant); RACT (Reactant or reagent)
 (bis- and mixed-tetrahydroisoquinolinium chlorofumarates as
 ultra-short-acting nondepolarizing neuromuscular blockers)

RN 213999-53-2 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2R)- (CA INDEX NAME)

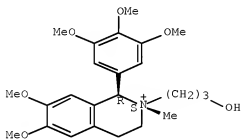
Absolute stereochemistry.

● Cl⁻

RN 220408-26-4 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

● Cl⁻

OSC.G 20 THERE ARE 20 CAPLUS RECORDS THAT CITE THIS RECORD (20 CITINGS)

RE.CNT 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 18 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1998:672522 CAPLUS Full-text

DN 129:275845

OREF 129:56249a

TI Substituted isoquinolines as ultra short acting neuromuscular blockers

IN Bigham, Eric Cleveland; Boswell, Grady Evan; Savarese, John Joseph;

Swaringen, Roy Archibald, Jr.; Patel, Sanjay Shashikant; Boros, Eric

Eugene; Mook, Robert Anthony, Jr.; Samano, Vincente

PA Glaxo Group Limited, UK; Cornell Research Foundation Inc.

SO PCT Int. Appl., 110 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 2

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|------------|------|----------|-----------------|--------------|
| PI | WO 9842675 | A1 | 19981001 | WO 1998-EP1652 | 19980323 <-- |

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RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, CH, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG

AU 9867305 A 19981020 AU 1998-67305 19980323 <--
 ZA 9802452 A 19990923 ZA 1998-2452 19980323 <--
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 EP 971898 B1 20050511

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 JP 2002503220 T 20020129 JP 1998-543233 19980323 <--
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 CZ 293786 B6 20040714 CZ 1999-3404 19980323
 EP 1526130 A1 20050427 EP 2005-1475 19980323

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 ES 2242275 T3 20051101 ES 1998-912494 19980323
 IN 1998CA00478 A 20051202 IN 1998-CA478 19980323
 AT 430734 T 20090515 AT 2003-20565 19980323
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FRAI GB 1997-6117 A 19970325
 GB 1997-24987 A 19971127
 EP 1998-912494 A3 19980323
 EP 1998-922626 A3 19980323
 JP 1998-543233 A3 19980323
 WO 1998-EP1652 W 19980323

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

OS MARPAT 129:275845

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Ultra short acting neuromuscular blocking agents I [q, t = 0-4; X1, X2 = halo; ha, hb = 0-2; Z1, Z2 = H, C1-6 alkyl, C2-6 alkenyl or C2-6 alkynyl with the proviso that Z1 and Z2 are not both hydrogen; Y1, Y2, Y3, Y4 = H, halo, C1-3 alkoxy; m, p = 1-6; n, r = 0-4; with the proviso that if ha and hb are both 0, then r is 0 and n is 0 to 2; R1-R14 = H, halo, C1-3 alkoxy, or R2 and R3 together with the carbon atoms to which they are bonded, R5 and R6 together with the carbon atoms to which they are bonded, R9 and R10 together with the carbon atoms to which they are bonded, R12 and R13 together with the carbon atoms to which they are bonded, may independently form a methylenedioxy or

ethylenedioxy moiety contained in a five- or six-membered ring; W1, W2 = C; A is a pharmaceutically acceptable anion], which are useful as skeletal muscle relaxants during emergency intubation procedures, routine surgery and post-operative settings, are disclosed. E.g., (Z)-2-chloro-1-[3-[(1S,2R)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-1,2,3,4-tetrahydro-2-isoquinolinio]propyl]-4-[3-[(1R,2S)-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-1,2,3,4-tetrahydro-2-isoquinolinio]propyl]-2-butenedioate dichloride was prepared

| | | | |
|----|--------------|--------------|--------------|
| IT | 213998-15-9P | 213998-16-0P | 213998-47-1P |
| | 213998-48-2P | 213998-49-3P | 213998-50-6P |
| | 213998-51-7P | 213998-52-8P | 213998-53-9P |
| | 213998-54-0P | 213998-55-1P | 213998-57-3P |
| | 213998-58-4P | 213998-59-5P | 213998-62-0P |
| | 213998-63-1P | 213998-64-2P | 213998-65-3P |
| | 213998-66-4P | 213998-67-5P | 213998-68-6P |
| | 213998-69-7P | 213998-70-0P | 213998-71-1P |
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| | 213999-31-6P | 213999-32-7P | 213999-38-3P |
| | 213999-39-1P | 213999-40-7P | 213999-41-8P |
| | 213999-42-9P | 213999-46-3P | 213999-47-4P |
| | 213999-50-9P | 213999-51-0P | 213999-52-1P |

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of substituted isoquinolines as ultra short acting neuromuscular blockers)

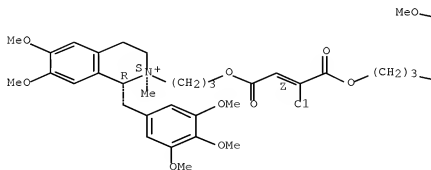
RN 213998-45-9 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

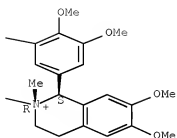
Double bond geometry as shown.

PAGE 1-A



● Cl⁻

PAGE 1-B

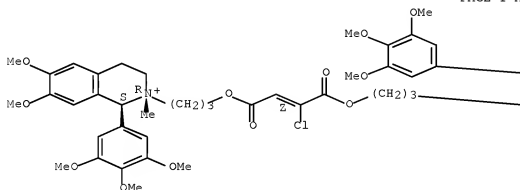


RN 213998-46-0 CAPLUS

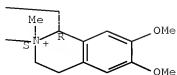
CN Isoquinolinium, 2-[3-[(1Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinol]propyl]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2). (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A



●2 C1-



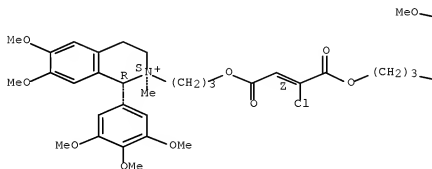
RN 213998-47-1 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

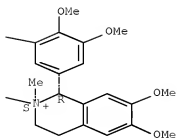
Double bond geometry as shown.

PAGE 1-A



● 2 Cl⁻

PAGE 1-B



RN 213998-48-2 CAPLUS

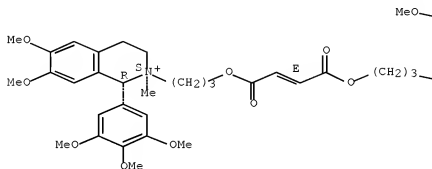
CN Isoquinolinium, 2,2'-[[(2E)-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-

10/591,174

propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

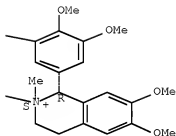
Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A



● 2 Cl-

PAGE 1-B

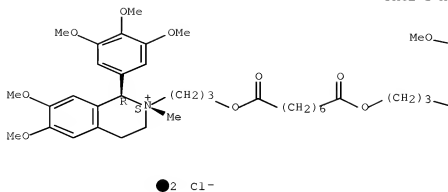


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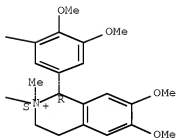
CN Isoquinolinium, 2,2'-[(1,8-dioxo-1,8-octanediyl)]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 1-B

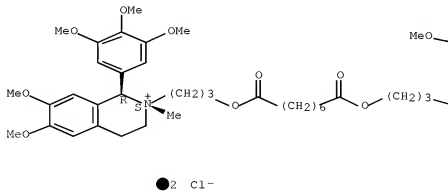


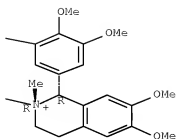
RN 213998-50-6 CAPLUS

CN Isoquinolinium, 2,2'-[(1,8-dioxo-1,8-octanediyloxybis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2R,2'S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

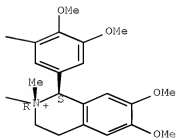
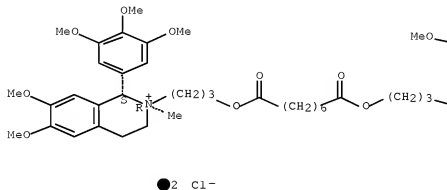




RN 213998-51-7 CAPLUS

CN Isoquinolinium, 2,2'-[(1,8-dioxo-1,8-octanediyl)bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,1'S,2R,2'R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

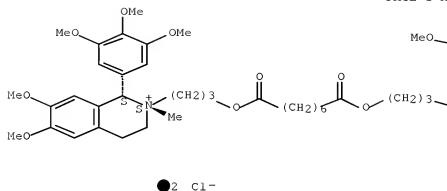


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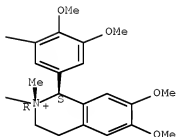
CN Isoquinolinium, 2,2'-[(1,8-dioxo-1,8-octanediyl)bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,1'S,2R,2'R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 1-B



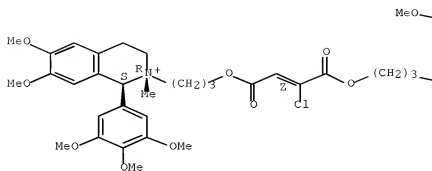
RN 213998-53-9 CAPLUS

CN Isoquinolinium, 2,2'-[[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,1'S,2R,2'R)- (9CI) (CA INDEX NAME)

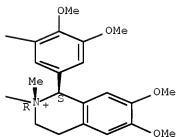
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

PAGE 1-B

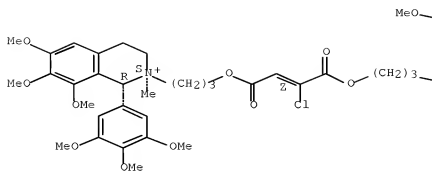
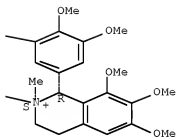


RN 213998-54-0 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1R,1'R,2S,2'S)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

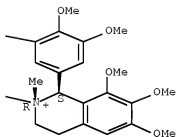
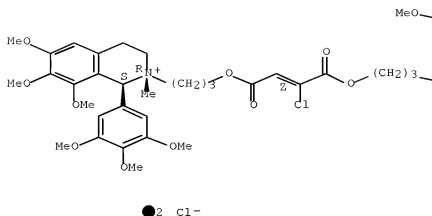
● 2 Cl⁻

RN 213998-55-1 CAPLUS

CN Isoquinolinium, 2,2'-[[(2Z)-2-chloro-1,4-dioxo-2-butene-1,4-diyl]bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,1'S,2R,2'R)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

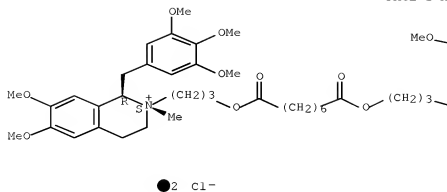


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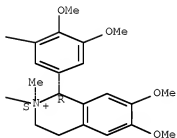
CN Isoquinolinium, 2-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A



PAGE 1-B

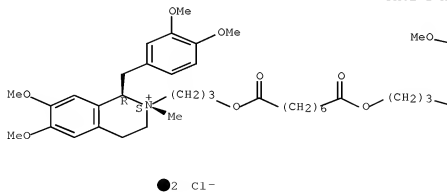


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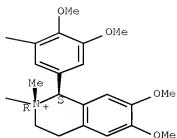
CN Isoquinolinium, 1-[(3,4-dimethoxyphenyl)methyl]-2-[3-[[1,8-dioxo-8-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]octyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 1-B

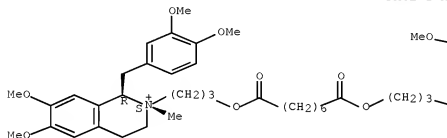


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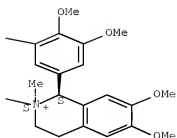
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Absolute stereochemistry.

PAGE 1-A

● 2 Cl⁻

PAGE 1-B



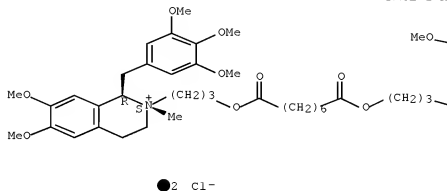
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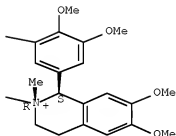
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Absolute stereochemistry.

PAGE 1-A



PAGE 1-B

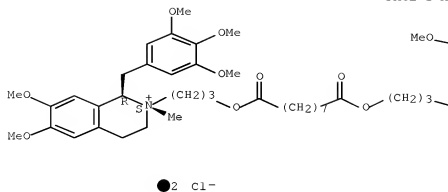


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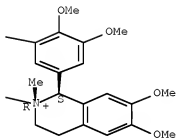
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Absolute stereochemistry.

PAGE 1-A



PAGE 1-B

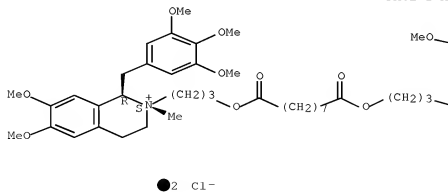


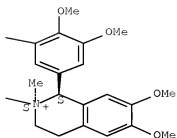
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Absolute stereochemistry.

PAGE 1-A

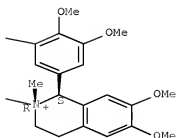
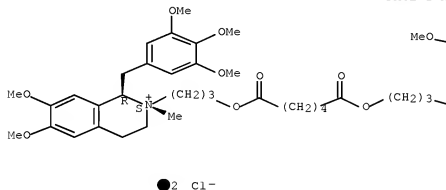




RN 213998-67-5 CAPLUS

CN Isoquinolinium, 2-[3-[[1,6-dioxo-6-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]hexyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



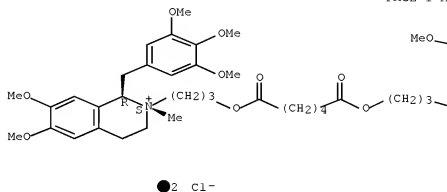
RN 213998-68-6 CAPLUS

CN Isoquinolinium, 2-[3-[[1,6-dioxo-6-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-

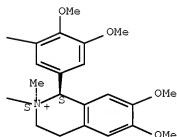
dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]hexyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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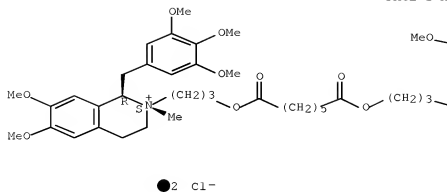


RN 213998-69-7 CAPLUS

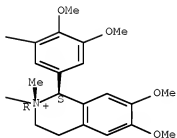
CN Isoquinolinium, 2-[3-[[1,7-dioxo-7-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]heptyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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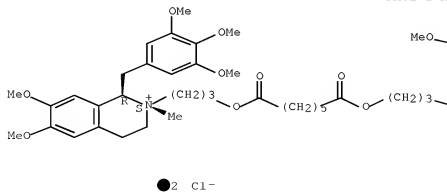


RN 213998-70-0 CAPLUS

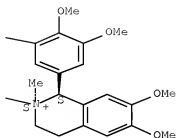
CN Isoquinolinium, 2-[3-[[1,7-dioxo-7-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]heptyloxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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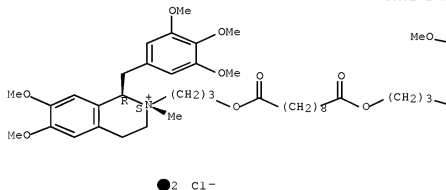


RN 213998-71-1 CAPLUS

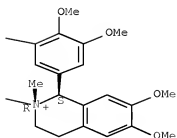
CN Isoquinolinium, 2-[3-[[1,10-dioxo-10-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]decyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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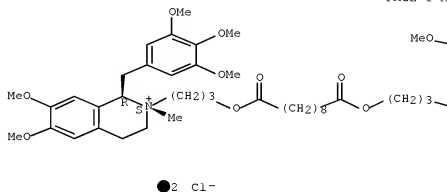
RN 213998-72-2 CAPLUS

CN Isoquinolinium, 2-[3-[[1,10-dioxo-10-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-

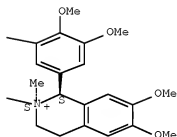
dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio[propoxy]decyl[oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

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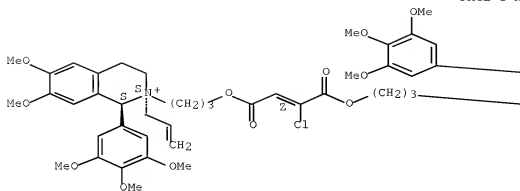
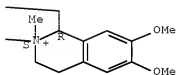


RN 213998-75-5 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]isoquinolinio[propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-(2-propenyl)-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

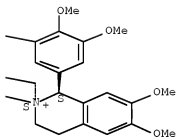
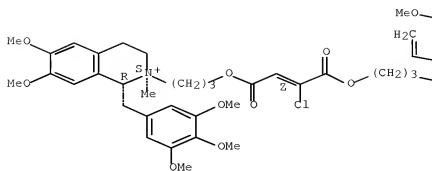
Double bond geometry as shown.

● 2 Cl⁻

RN 213998-76-6 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-(2-propenyl)-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,2S)- (9CI) (CA INDEX NAME)

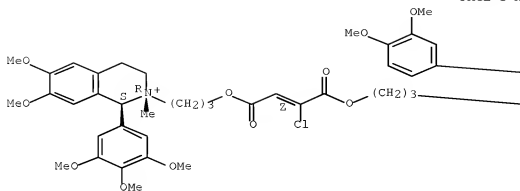
Absolute stereochemistry.
Double bond geometry as shown.



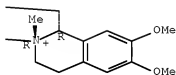
RN 213998-77-7 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-3-chloro-4-[3-[(1R,2R)-1-[(3,4-dimethoxyphenyl)methyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-1-methylisoquinolinio]propoxy]-1,4-dioxo-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.



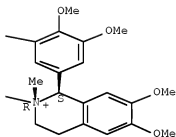
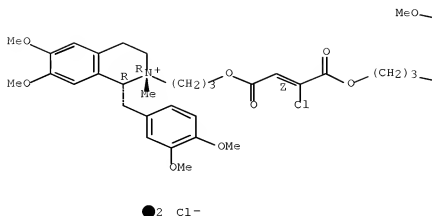
● 2 Cl-



RN 213998-78-8 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-4-[3-[(1R,2R)-1-[(3,4-dimethoxyphenyl)methyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methylisoquinolinio]propoxy]-1,4-dioxo-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,2R)- (9CI) (CA INDEX NAME)

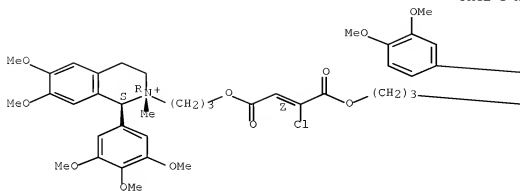
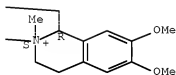
Absolute stereochemistry.
Double bond geometry as shown.



RN 213998-79-9 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-4-[3-[(1R,2S)-1-[(3,4-dimethoxyphenyl)methyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-1-methylisoquinolinio]propoxy]-1,4-dioxo-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

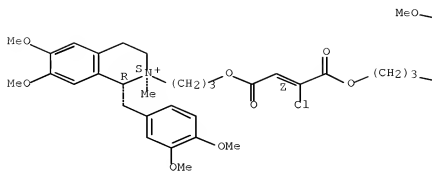
●2 Cl⁻

RN 213998-80-2 CAPLUS

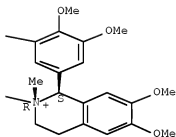
CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-4-[3-[(1R,2S)-1-[(3,4-dimethoxyphenyl)methyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methylisoquinolinio]propoxy]-1,4-dioxo-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, (1S,2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

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● 2 Cl⁻

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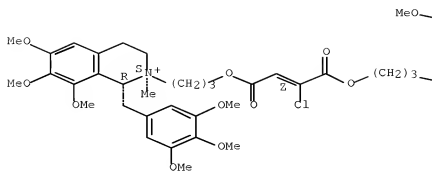


RN 213998-81-3 CAPLUS

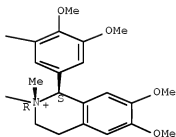
CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)]

Absolute stereochemistry.
Double bond geometry as shown.

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● 2 Cl⁻

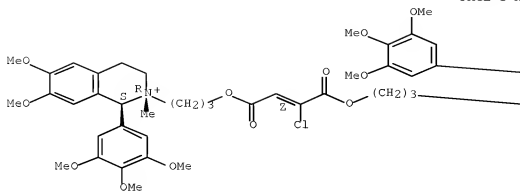
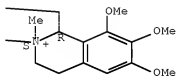
PAGE 1-B



RN 213998-82-4 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

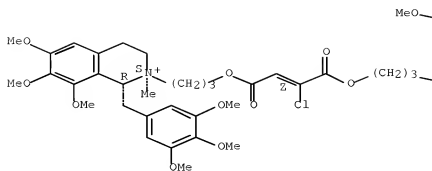
RN 213998-83-5 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)]

Absolute stereochemistry.

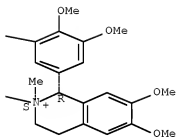
Double bond geometry as shown.

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● 2 Cl-

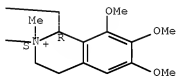
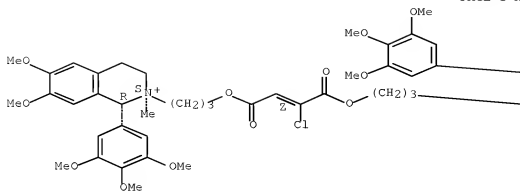
PAGE 1-B



RN 213998-84-6 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

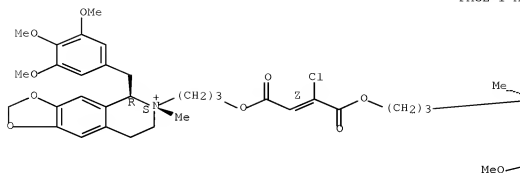


RN 213999-19-0 CAPLUS

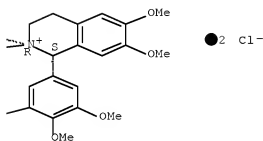
CN 1,3-Dioxolo[4,5-g]isoquinolinium, 6-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-buten-1-yl]oxy]propyl]-5,6,7,8-tetrahydro-6-methyl-5-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (5R,6S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

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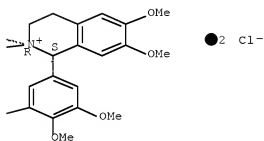
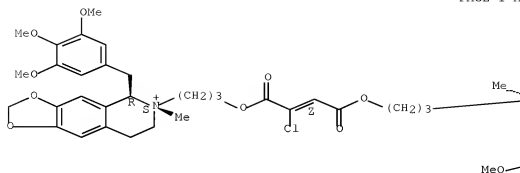


RN 213999-20-3 CAPLUS

CN 1,3-Dioxolo[4,5-g]isoquinolinium, 6-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-buten-1-yloxy]propyl]-5,6,7,8-tetrahydro-6-methyl-5-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (5R,6S)- (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

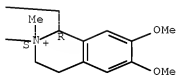
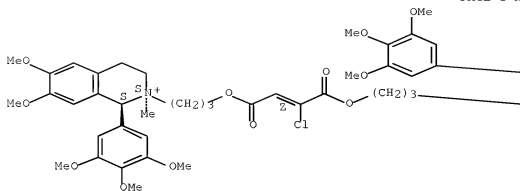


RN 213999-24-7 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

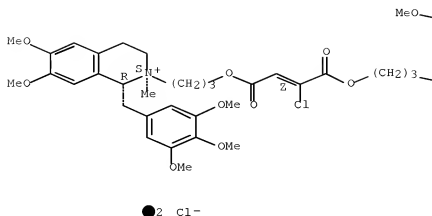


RN 213999-25-8 CAPLUS

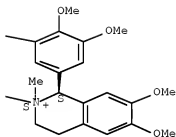
CN Isoquinolinium, 2-[3-[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

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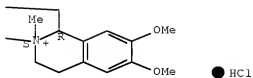
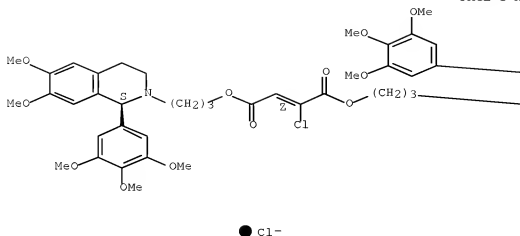
PAGE 1-B



RN 213999-26-9 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-4-[3-[(1S)-3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-2(1H)-isoquinolinyl]propoxy]-1,4-dioxo-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride, hydrochloride (1:1:1), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

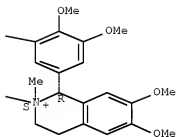
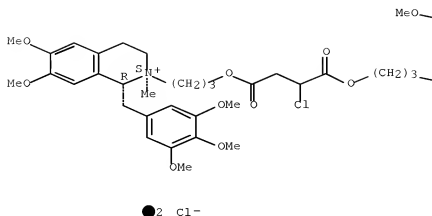


RN 213999-27-0 CAPLUS

CN Isoquinolinium, 2-[3-[(2E)-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

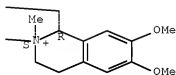
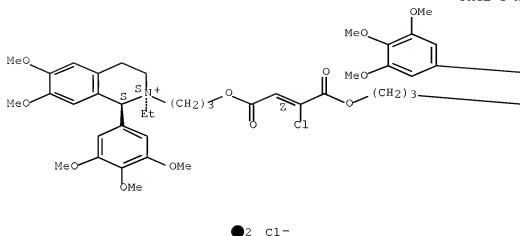
Double bond geometry as shown.



RN 213999-31-6 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-4-[3-[(1S,2S)-2-ethyl-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-1,4-dioxo-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

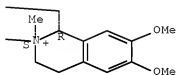
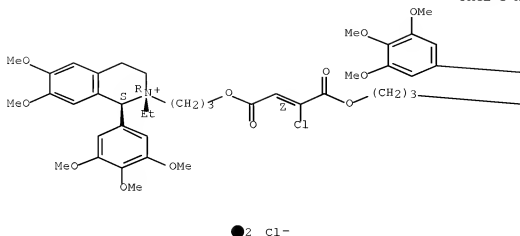
Absolute stereochemistry.
Double bond geometry as shown.



RN 213999-32-7 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-4-[3-[(1S,2R)-2-ethyl-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-1,4-dioxo-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

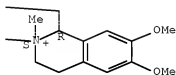
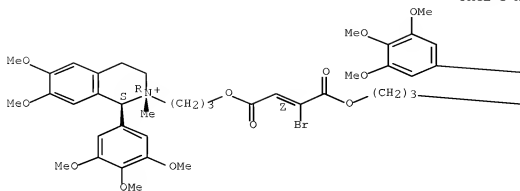
Absolute stereochemistry.
Double bond geometry as shown.



RN 213999-38-3 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-bromo-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]propyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dibromide, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

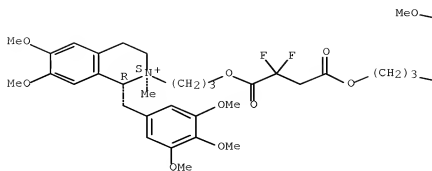


RN 213999-39-4 CAPLUS

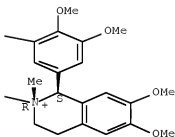
CN Isoquinolinium, 2-[3-[2,2-difluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

● 2 Cl⁻

PAGE 1-B

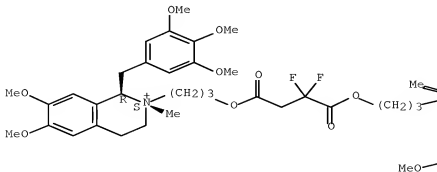


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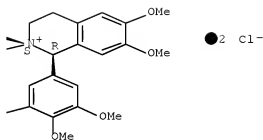
CN Isoquinolinium, 2-[3-[3,3-difluoro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A



PAGE 1-B

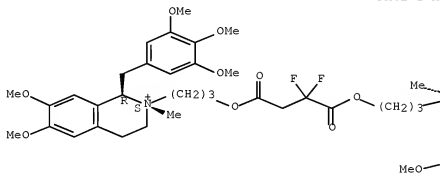


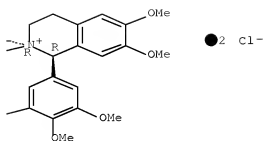
RN 213999-41-8 CAPLUS

CN Isoquinolinium, 2-[3-[3,3-difluoro-1,4-dioxo-4-[3-[(1R,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

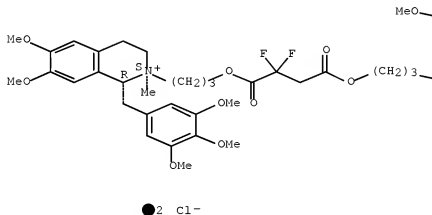


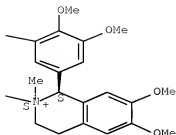


RN 213999-42-9 CAPLUS

CN Isoquinolinium, 2-[3-[2,2-difluoro-1,4-dioxo-4-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

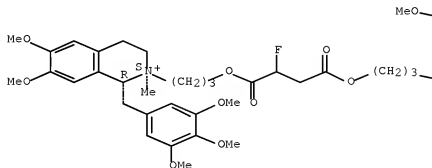
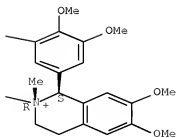




RN 213999-46-3 CAPLUS

CN Isoquinolinium, 2-[3-[2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

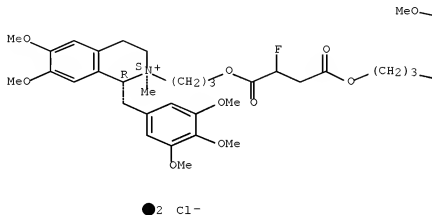
● 2 Cl⁻

RN 213999-47-4 CAPLUS

CN Isoquinolinium, 2-[3-[2-fluoro-1,4-dioxo-4-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy)butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

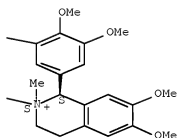
Absolute stereochemistry.

PAGE 1-A



● 2 Cl⁻

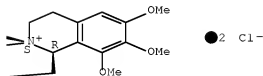
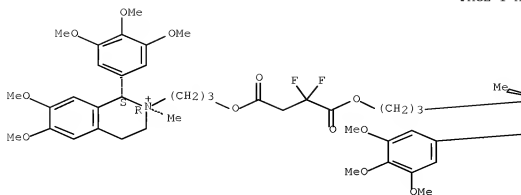
PAGE 1-B



RN 213999-50-9 CAPLUS

CN Isoquinolinium, 2-[3-[2,2-difluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy)butoxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

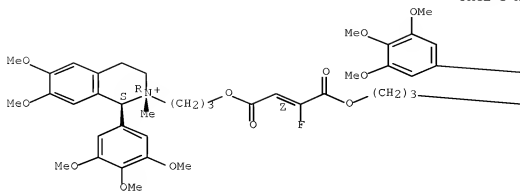


RN 213999-51-0 CAPLUS

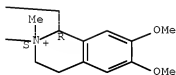
CN Isoquinolinium, 2-[3-[[(2Z)-2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.



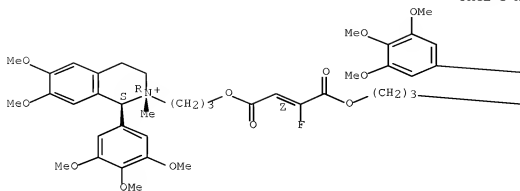
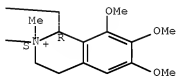
● 2 Cl⁻



RN 213999-52-1 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

IT 22325-16-2

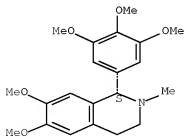
RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of substituted isoquinolines as ultra short acting neuromuscular blockers)

RN 22325-16-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



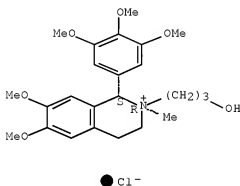
IT 213999-53-2P 213999-54-3P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation of substituted isoquinolines as ultra short acting neuromuscular blockers)

RN 213999-53-2 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2R)- (CA INDEX NAME)

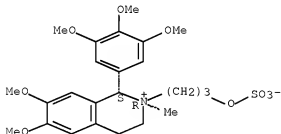
Absolute stereochemistry.



RN 213999-54-3 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-2-[3-(sulfooxy)propyl]-1-(3,4,5-trimethoxyphenyl)-, inner salt, (1S,2R)- (CA INDEX NAME)

Absolute stereochemistry.



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
 RE.CNT 11 THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 19 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1998:672521 CAPLUS [Full-text](#)

DN 129:290071

OREF 129:59118h,59119a

TI Preparation of dimeric isoquinolines as ultra short acting neuromuscular blockers.

IN Bigham, Eric Cleveland; Boswell, Grady Evan; Savarese, John Joseph; Swaringen, Roy Archibald, Jr.; Patel, Sanjay Shashikant; Boros, Eric Eugene; Mook, Robert Anthony, Jr.; Samano, Vincente

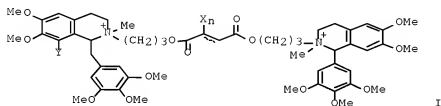
PA Glaxo Group Limited, UK; Cornell Research Foundation Inc.

SO PCT Int. Appl., 49 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 2

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| | EE 9900424 | A | 20000417 | EE 1999-424 | 19980323 <-- |
| | EE 4113 | B1 | 20030815 | | |
| | BR 9808422 | A | 20000523 | BR 1998-8422 | 19980323 <-- |
| | NZ 337797 | A | 20010330 | NZ 1998-337797 | 19980323 <-- |
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| | EP 1380573 | A2 | 20040114 | EP 2003-20565 | 19980323 <-- |
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| | CN 1203061 | C | 20050525 | CN 1998-805300 | 19980323 |
| | IL 131918 | A | 20050925 | IL 1998-131918 | 19980323 |
| | ES 2242275 | T3 | 20051101 | ES 1998-912494 | 19980323 |
| | IN 1998CA00478 | A | 20051202 | IN 1998-CA478 | 19980323 |
| | PL 190860 | B1 | 20060228 | PL 1998-335885 | 19980323 |
| | AP 1797 | A | 20071231 | AP 1999-1658 | 19980323 |
| | AT 430734 | T | 20090515 | AT 2003-20565 | 19980323 |
| | HR 9800157 | B1 | 20041031 | HR 1998-157 | 19980324 |
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| | GB 1997-24987 | A | 19971127 |
| | EP 1998-912494 | A3 | 19980323 |
| | EP 1998-922626 | A3 | 19980323 |
| | JP 1998-543233 | A3 | 19980323 |
| | WO 1998-EP1651 | W | 19980323 |

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT
 OS MARPAT 129:290071
 GI



AB Title compds. (I; X = halo; n = 1, 2; Y = H, OMe; A = pharmaceutically acceptable anion), were prepared Thus, 2,2-difluoro-4-[3-[(1S,2R)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-1,2,3,4-tetrahydro-2-isoquinolinio]propyl]-1-[3-[(1R,2S)-2-methyl-6,7,8-trimethoxy-1-(3,4,5-trimethoxyphenyl)methyl]-1,2,3,4-tetrahydro-2-isoquinolinio]propyl]butanedioate dichloride showed ED₉₅ = 0.035 mg/kg i.v. for blockage of evoked twitch response of the tibialis anterior muscle.

| | | | |
|----|--------------|--------------|--------------|
| IT | 213996-45-9P | 213998-46-0P | 213998-57-3P |
| | 213996-58-4P | 213998-81-2P | 213998-62-4P |
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| | 213999-52-1P | | |

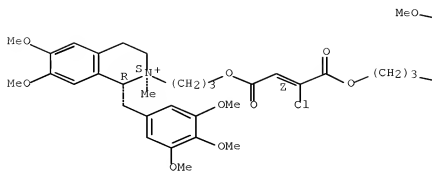
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of dimeric isoquinolines as ultra short acting neuromuscular blockers)

RN 213998-45-9 CAPLUS

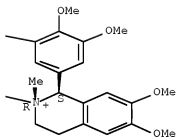
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Absolute stereochemistry.
 Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

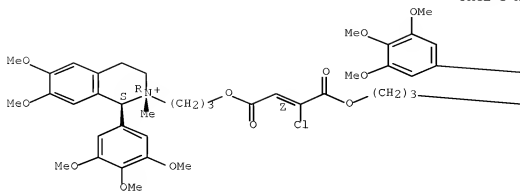
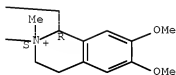
PAGE 1-B



RN 213998-46-0 CAPLUS

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Absolute stereochemistry.
Double bond geometry as shown.

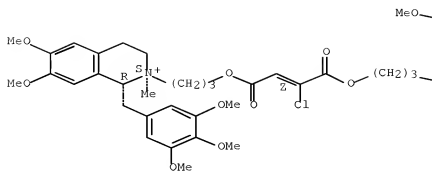
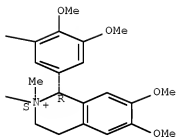
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RN 213998-57-3 CAPLUS

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Absolute stereochemistry.

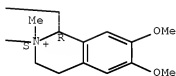
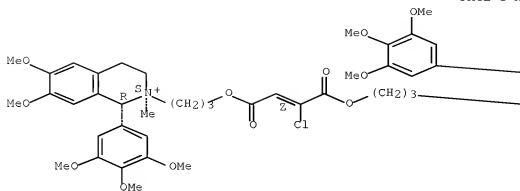
Double bond geometry as shown.

● 2 Cl⁻

RN 213998-58-4 CAPLUS

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Absolute stereochemistry.
Double bond geometry as shown.

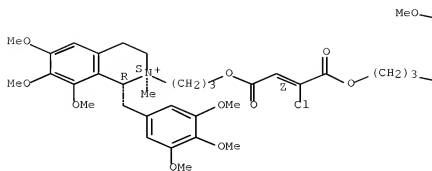


RN 213998-81-3 CAPLUS

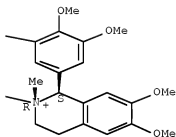
CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

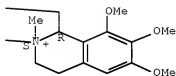
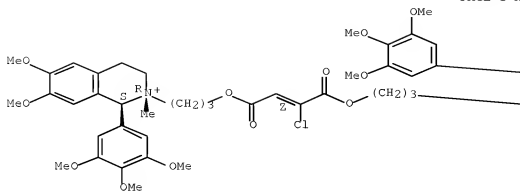
PAGE 1-B



RN 213998-82-4 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

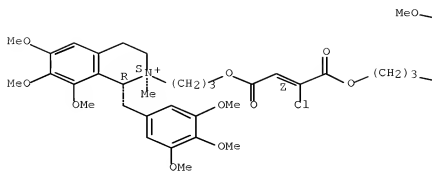


RN 213998-83-5 CAPLUS

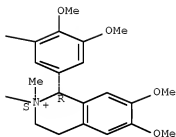
CN Isoquinolinium, 2-[3-[[[(2Z)-3-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
Double bond geometry as shown.

PAGE 1-A

● 2 Cl⁻

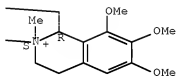
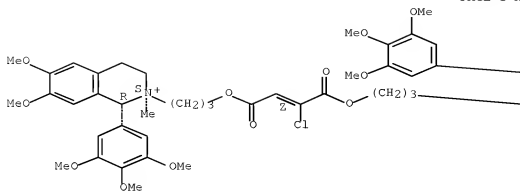
PAGE 1-B



RN 213998-84-6 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1R,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]-2-buten-1-yl]oxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

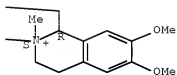
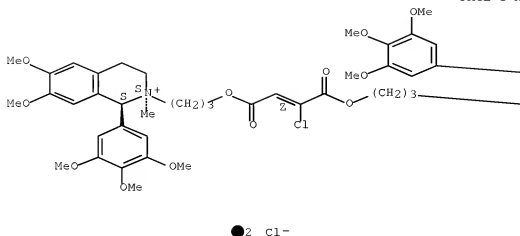
Absolute stereochemistry.
Double bond geometry as shown.



RN 213999-24-7 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

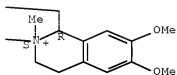
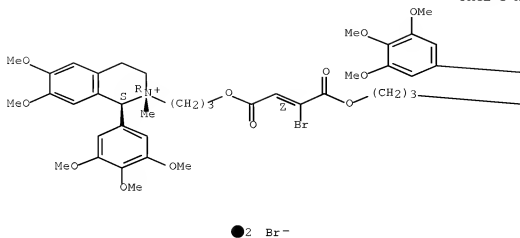
Absolute stereochemistry.
Double bond geometry as shown.



RN 213999-38-3 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-bromo-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]propyl]-2-butenyl]oxypropyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dibromide, (1R,2S)- (9CI) (CA INDEX NAME)

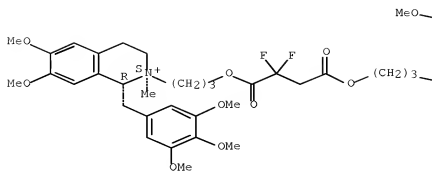
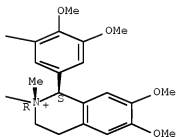
Absolute stereochemistry.
Double bond geometry as shown.



RN 213999-39-4 CAPLUS

CN Isoquinolinium, 2-[3-[2,2-difluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

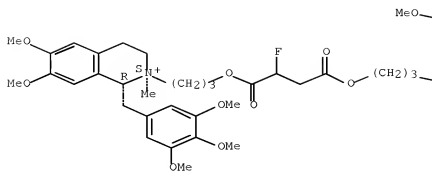
● 2 Cl⁻

RN 213999-46-3 CAPLUS

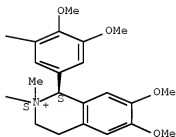
CN Isoquinolinium, 2-[3-[2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

● 2 Cl⁻

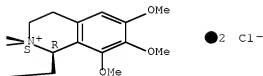
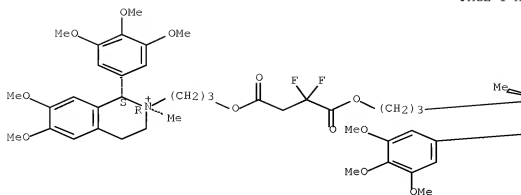
PAGE 1-B



RN 213999-50-9 CAPLUS

CN Isoquinolinium, 2-[3-[2,2-difluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinium-2-yl]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

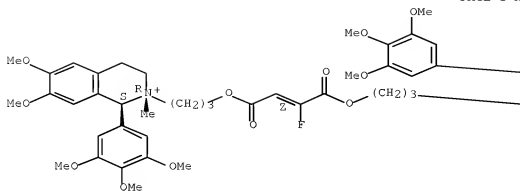
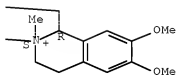


RN 213999-51-0 CAPLUS

CN Isoquinolinium, 2-[3-[[(2Z)-2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

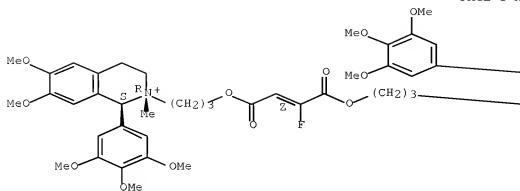
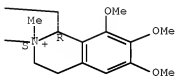
Double bond geometry as shown.

● 2 Cl⁻

RN 213999-52-1 CAPLUS

CN Isoquinolinium, 2-[3-[[[(2Z)-2-fluoro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]propoxy]-2-buten-1-yl]oxypropyl]-1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)]

Absolute stereochemistry.
Double bond geometry as shown.

● 2 Cl⁻

IT 22325-16-2

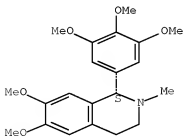
RL: RCT (Reactant); RACT (Reactant or reagent)

(preparation of dimeric isoquinolines as ultra short acting neuromuscular blockers)

RN 22325-16-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



IT 213999-42-9P 213999-53-2P 213999-54-3P

14191-49-8P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

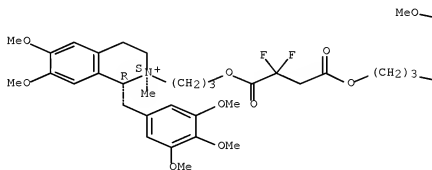
(preparation of dimeric isoquinolines as ultra short acting neuromuscular blockers)

RN 213999-42-9 CAPLUS

CN Isoquinolinium, 2-[3-[2,2-difluoro-1,4-dioxo-4-[3-[(1S,2S)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]butoxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, dichloride, (1R,2S)- (9CI) (CA INDEX NAME)

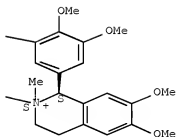
Absolute stereochemistry.

PAGE 1-A



● 2 Cl⁻

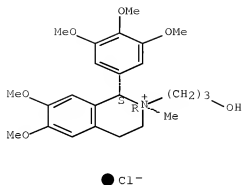
PAGE 1-B



RN 213999-53-2 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2R)- (CA INDEX NAME)

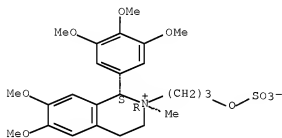
Absolute stereochemistry.



RN 213999-54-3 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-2-[3-(sulfooxy)propyl]-1-(3,4,5-trimethoxyphenyl)-, inner salt, (1S,2R)- (CA INDEX NAME)

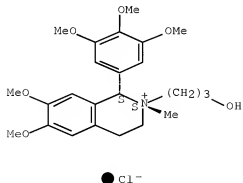
Absolute stereochemistry.



RN 214191-49-8 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1S,2S)- (CA INDEX NAME)

Absolute stereochemistry.



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
 RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 20 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1997:661349 CAPLUS Full-text

DN 127:341647

OREF 127:66922h,66923a

TI Neuromuscular blocking activity of cyclic and acyclic bis-quaternary ammonium analogs of mivacurium chloride in the cat

AU Patel, Sanjay S.; Maehr, R.; Savarese, John J.; Jackson, Mary M.; Wastila, William B.; Wisowaty, James C.

CS Chem. Dev. Lab., Burroughs Wellcome Co., Research Triangle Park, NC, 27709, USA

SO European Journal of Pharmaceutical Sciences (1997), 5(5), 253-266

CODEN: EPSCED; ISSN: 0928-0987

PB Elsevier

DT Journal

LA English

AB The purpose of this work was to identify a new ultra-short-acting neuromuscular blocking agent devoid of the potential to produce cardiovascular effects at \geq ED95 doses. Four new bis-quaternary mivacurium analogs that are acyclic with respect to the bis-isoquinolinium nuclei and seven new bis-quaternary mivacurium analogs that are derivs. of (E)-oct-4-enedioic acid, (E)-oct-2-enedioic acid, and (E)-oct-4-enedithioic acid, were synthesized and tested for neuromuscular blocking activity in the cat. In general, compared with mivacurium, the acyclic analogs were of much lower potency but showed a faster onset (time from injection to maximum neuromuscular block) and a much shorter duration of action (time from injection to 95% recovery) at approx. ED95 doses. However, these acyclic analogs had a considerably narrower safety margin (i.e., the ratio of doses that produce unwanted cardiovascular or autonomic effects to those that produce neuromuscular block) than mivacurium. The (E)-oct-4-enedioate and (E)-oct-4-enedithioate analogs showed a neuromuscular blocking profile similar to the acyclic analogs. The (E)-oct-2-enedioate isomer of mivacurium did not have any advantageous neuromuscular blocking properties over mivacurium and, in fact, elicited cardiovascular and autonomic effects at much lower multiples of ED95. Structural changes to mivacurium, however minor, to either the inter-onium chain or the onium centers (or both) result in compds. whose cardiovascular and autonomic safety profiles are highly compromised in return for the desirable rapid onset and brevity of neuromuscular blocking action at \geq ED95 doses. The intact isoquinolinium nucleus appears to confer a superior safety profile over that of an acyclic onium nucleus.

IT 198400-96-3F

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(neuromuscular blocking activity of cyclic and acyclic bis-quaternary ammonium analogs of mivacurium chloride)

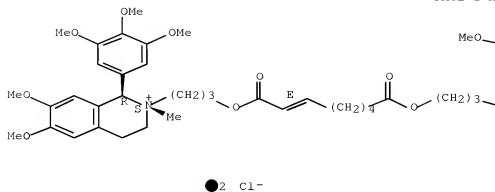
RN 198400-96-3 CAPLUS

CN Isoquinolinium, 2,2'-[(1,8-dioxo-2-octene-1,8-diyl)bis(oxy-3,1-propanediyl)]bis[1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, dichloride, [1a,2β[E(1'R*,2'S*)]]- (9CI)
(CA INDEX NAME)

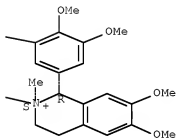
Relative stereochemistry.

Double bond geometry as shown.

PAGE 1-A



PAGE 1-B



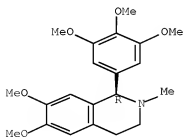
IT 33033-86-2

RL: RCT (Reactant); RACT (Reactant or reagent)
(neuromuscular blocking activity of cyclic and acyclic bis-quaternary ammonium analogs of mivacurium chloride)

RN 33033-86-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



IT 198401-05-7P

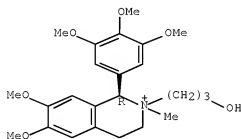
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(neuromuscular blocking activity of cyclic and acyclic bis-quaternary

ammonium analogs of mivacurium chloride)

RN 198401-05-7 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-2-(3-hydroxypropyl)-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, chloride (1:1), (1R)- (CA INDEX NAME)

Absolute stereochemistry.

● Cl⁻

OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)
 RE.CNT 22 THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 21 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1997:183784 CAPLUS Full-text

DN 126:233107

OREF 126:44929a,44932a

TI An application of TLC chromatographic data in QSAR assay of TIQs derivatives with β 2-adrenergic activity. Part I.

AU Brzezinska, Elzbieta

CS Institute of Chemistry and Technology of Drugs, School of Medicine, Lodz, 90-151, Pol.

SO Acta Poloniae Pharmaceutica (1996), 53(5), 383-388

CODEN: APPHAX; ISSN: 0001-6837

PB Polish Pharmaceutical Society

DT Journal

LA English

AB A QSAR anal. of β 2-adrenergic activity and chromatog. data of 4,6,8-trihydroxy-, 6,7-dihydroxy- and 6,7-dimethoxy-1,2,3,4-tetrahydroisoquinoline derivs. were made. The TLC plates (silica gel 60 F254 silanized precoated), impregnated with solns. of selected amino acids mixts., were used as a β 2-agonistic and antagonistic interaction models. The hydrophobicity data of examined compds. (π and Σf -values) were obtained and used in the QSAR assay. Using a linear regression anal., interrelations between chromatog. and biol. activity data were found.

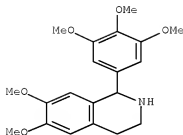
IT 33633-84-0 199553-85-7

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

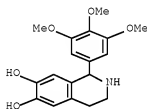
(QSAR anal. of β 2-adrenergic activity and chromatog. data of trihydroxy-, dihydroxy- and dimethoxy-tetrahydroisoquinoline derivs.)

RN 33033-84-0 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)

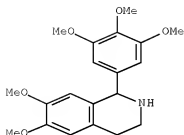


RN 188553-85-7 CAPLUS
 CN 6,7-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)

L5 ANSWER 22 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1997:183781 CAPLUS [Full-text](#)
 DN 126:287916
 OREF 126:55561a,55564a
 TI Synthesis and pharmacological properties of
 6,7-dimethoxy-1,2,3,4-tetrahydroisoquinoline derivatives
 AU Brzezinska, Elzbieta
 CS Institute of Chemistry and Technology of Drugs, School of Medicine, Lodz,
 90-151, Pol.
 SO Acta Poloniae Pharmaceutica (1996), 53(5), 365-371
 CODEN: APPHAX; ISSN: 0001-6837
 PB Polish Pharmaceutical Society
 DT Journal
 LA English
 AB Some selected 1-aryl-6,7-dimethoxy-1,2,3,4-tetrahydroisoquinoline derivs. were
 synthesized and evaluated as the β -adrenoceptor agents. Some of the compds.
 showed a weak agonistic or antagonistic activity on these receptors.
 IT 33033-84-0F
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); SPN (Synthetic preparation); BIOL (Biological
 study); PREP (Preparation)
 (synthesis and pharmacol. properties of
 6,7-dimethoxy-1,2,3,4-tetrahydroisoquinoline derivs.)
 RN 33033-84-0 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
 (CA INDEX NAME)



OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)

L5 ANSWER 23 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1996:376527 CAPLUS Full-text

DN 125:167761

OREF 125:31433a

TI Asymmetric synthesis. XLI. Totally stereoselective synthesis of 1,3-disubstituted tetrahydroisoquinolines via the CN(R,S) method

AU Gosmann, Grace; Guillaume, Dominique; Husson, Henri-Philippe

CS Lab. Chimie Therapeutique associe CNRS, Univ. R. Descartes, Paris, 75270, Fr.

SO Tetrahedron Letters (1996), 37(25), 4369-4372

CODEN: TELEAY; ISSN: 0040-4039

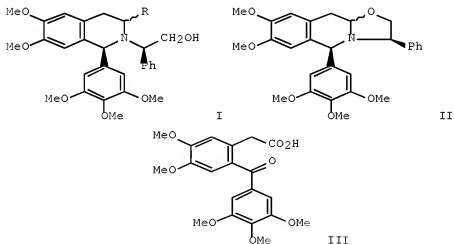
PB Elsevier

DT Journal

LA English

OS CASREACT 125:167761

GI



AB Optically active cis- or trans- 1,3-disubstituted tetrahydroisoquinolines 3-R- or 3S-I (R = Me, CH₂Ph) can be prepared selectively from the same oxazolidine II. This latter is easily obtained from keto-acid III and (R)-(-)phenylglycinol.

IT 180072-45-1P 180072-48-4P 180072-49-5P
180072-50-8P

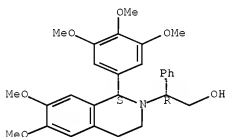
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)

(stereoselective preparation of isoquinolines)

RN 180072-45-1 CAPLUS

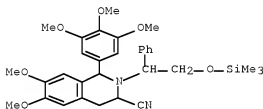
CN 2-(1H)-Isoquinolineethanol, 3,4-dihydro-6,7-dimethoxy- β -phenyl-1-
(3,4,5-trimethoxyphenyl)-, [S-(R*,S*)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



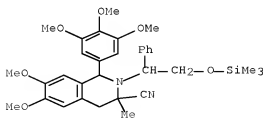
RN 180072-48-4 CAPLUS

CN 3-Isoquinolinecarbonitrile, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-[(1R)-1-
phenyl-2-[(trimethylsilyl)oxy]ethyl]-1-(3,4,5-trimethoxyphenyl)-, (1S)-
(CA INDEX NAME)



RN 180072-49-5 CAPLUS

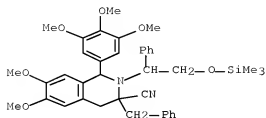
CN 3-Isoquinolinecarbonitrile, 1,2,3,4-tetrahydro-6,7-dimethoxy-3-methyl-2-
[(1R)-1-phenyl-2-[(trimethylsilyl)oxy]ethyl]-1-(3,4,5-trimethoxyphenyl)-,
(1S)- (CA INDEX NAME)



RN 180072-50-8 CAPLUS

CN 3-Isoquinolinecarbonitrile, 1,2,3,4-tetrahydro-6,7-dimethoxy-3-

(phenylmethyl)-2-[(1R)-1-phenyl-2-[(trimethylsilyl)oxy]ethyl]-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)



IT 32886-69-4P 180072-46-2P 180072-47-3P

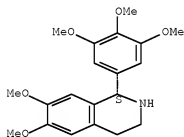
180187-56-8P 180187-57-9P

RL: SPN (Synthetic preparation); PREP (Preparation)
(stereoselective preparation of isoquinolines)

RN 32886-69-4 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, (S)- (9CI) (CA INDEX NAME)

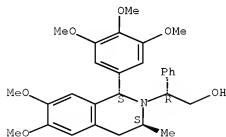
Absolute stereochemistry.



RN 180072-46-2 CAPLUS

CN 2-(1H)-Isoquinolineethanol, 3,4-dihydro-6,7-dimethoxy-3-methyl-β-phenyl-1-(3,4,5-trimethoxyphenyl)-, [1S-[1α,2(S*),3β]]- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.

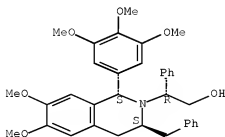


RN 180072-47-3 CAPLUS

CN 2-(1H)-Isoquinolineethanol, 3,4-dihydro-6,7-dimethoxy-β-phenyl-1-(3,4,5-trimethoxyphenyl)-, [1S-[1α,2(S*),3β]]- (9CI)
(CA INDEX NAME)

(phenylmethyl)-1-(3,4,5-trimethoxyphenyl)-, [1S-[1 α ,2(S*),3 β]]-
(9CI) (CA INDEX NAME)

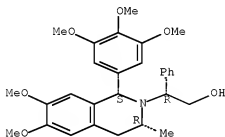
Absolute stereochemistry.



RN 180187-56-8 CAPLUS

CN 2(1H)-Isoquinolineethanol, 3,4-dihydro-6,7-dimethoxy-3-methyl-β-
phenyl-1-(3,4,5-trimethoxyphenyl)-, [1S-[1 α ,2(S*),3 α]]- (9CI)
(CA INDEX NAME)

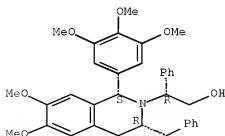
Absolute stereochemistry.



RN 180187-57-9 CAPLUS

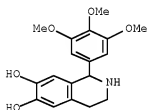
CN 2(1H)-Isoquinolineethanol, 3,4-dihydro-6,7-dimethoxy-β-phenyl-3-
(phenylmethyl)-1-(3,4,5-trimethoxyphenyl)-,
[1S-[1 α ,2(S*),3 α]]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



OSC.G 13 THERE ARE 13 CAPLUS RECORDS THAT CITE THIS RECORD (13 CITINGS)

L5 ANSWER 24 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1996:365418 CAPLUS [Full-text](#)
 DN 125:75373
 OREF 125:14059a,14062a
 TI Synthesis and pharmacological investigations of
 1,2,3,4-tetrahydroisoquinoline derivatives
 AU Brzezinska, E.; Venter, D.; Glinka, R.
 CS Institute Chemistry Technology Drugs, Medical University Lodz, Lodz,
 90-151, Pol.
 SO Pharmazie (1996), 51(6), 397-399
 CODEN: PHARAT; ISSN: 0031-7144
 PB Govi-Verlag Pharmazeutischer Verlag
 DT Journal
 LA English
 AB Selected 1-aryl-6,7-dihydroxy-1,2,3,4-tetrahydroisoquinoline derivs. were
 synthesized and evaluated to determination the role of the hydrophobic Ph
 group at C-1 on the activity of these derivs. at β -adrenoreceptors. All the
 tests show that the investigated compds. are not very active. The Ph group is
 a poor substituent for β -adrenomimetic activity, but the presence of the Ph
 substituent at position 1 appears to enable these mol. to act as antagonists
 at β -adrenoreceptors.
 IT 57529-51-8P
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
 BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of tetrahydroisoquinoline derivs. and aryl group effect on
 β -adrenoreceptors)
 RN 57529-51-8 CAPLUS
 CN 6,7-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)-,
 hydrochloride (1:1) (CA INDEX NAME)

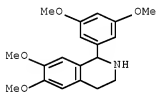


● HCl

OSC.G 6 THERE ARE 6 CAPLUS RECORDS THAT CITE THIS RECORD (6 CITINGS)

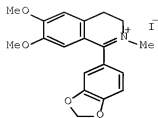
L5 ANSWER 25 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1996:60166 CAPLUS [Full-text](#)
 DN 124:202678
 OREF 124:37481a,37484a
 TI Synthesis of 3,4-dihydroisoquinolines,
 2-alkyl(acyl)-1(2H)-3,4-dihydroisoquinolinones,
 2-alkyl-1(2H)-isoquinolinones and 1-alkyl-2(2H)-quinolinones by oxidation
 with potassium permanganate
 AU Venkov, Atanas P.; Statkova-Abeghe, Stela M.
 CS Dep. Chem., Univ. Plovdiv, Plovdiv, 4000, Bulg.
 SO Tetrahedron (1996), 52(4), 1451-60
 CODEN: TETRAJ; ISSN: 0040-4020

PB Elsevier
 DT Journal
 LA English
 OS CASREACT 124:202678
 AB Synthesis of 3,4-dihydroisoquinolines, 2-alkyl- and 2-acyl-3,4-dihydro-1(2H)-isoquinolinones, 2-alkyl-1(2H)-isoquinolinones, N-alkyl-3,4-dihydro-2(2H)-quinolinones and N-alkyl-2(2H)-quinolinones by oxidation of 1,2,3,4-tetrahydroisoquinolines, N-alkyl (acyl)iminium salts of 3,4-dihydroisoquinolines and isoquinoline as well as of N-alkyl ammonium salts of tetrahydroquinoline and quinoline with potassium permanganate is described.
 IT 174503-32-3
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (synthesis of 3,4-dihydroisoquinolines,
 2-alkyl(acyl)-1(2H)-3,4-dihydroisoquinolinones,
 2-alkyl-1(2H)-isoquinolinones and 1-alkyl-2(2H)-quinolinones by oxidation
 with potassium permanganate)
 RN 174503-32-3 CAPLUS
 CN Isoquinoline, 1-(3,5-dimethoxyphenyl)-1,2,3,4-tetrahydro-6,7-dimethoxy-
 (CA INDEX NAME)

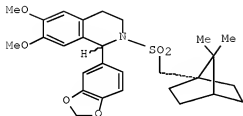


OSC.G 26 THERE ARE 26 CAPLUS RECORDS THAT CITE THIS RECORD (26 CITINGS)

L5 ANSWER 26 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1995:447155 CAPLUS Full-text
 DN 123:56331
 OREF 123:10159a,10162a
 TI Some observations on the enantio- and diastereo-selective synthesis of
 1-substituted-1,2,3,4-tetrahydroisoquinolines
 AU Nagarajan, K.; Chandrasekharan, J.; Rodrigues, P. J.
 CS R and D Centre, Searle India Ltd., Thane, 400 601, India
 SO Journal of the Indian Institute of Science (1994), 74(2), 247-55
 CODEN: JIISAD; ISSN: 0019-4964
 DT Journal
 LA English
 GI



I



II

AB Several approaches to the synthesis of optically active 1-aryl-1,2,3,4-tetrahydroisoquinoline alkaloids, cryptostylinines from dihydroisoquinoline precursors have been tried. Redns. of the 1-aryldihydroisoquinolines as well as of their methiodides with yeast are unsuccessful. Reduction of the quaternary salts, e.g. I, with sodium tris-acyloxyborohydrides gives the tetrahydroisoquinoline alkaloids in unsatisfactory enantiomeric excess. (\pm)-Norcryptostyline is resolved with (-)- and (+)-tartaric acid into (-)S and (+)R enantiomers, which are converted to their camphorsulfonyl derivs. II. Reaction of camphorsulfonyl homoveratryl amine with piperonal affords a mixture of 1R-II and 1S-II in the ratio of 4:3, whereas camphorsulfonylation of (\pm)-norcryptostyline gives 1R-II much in excess of 1S-II (4:1).

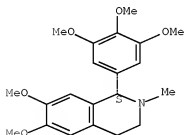
IT 22325-16-2P, Cryptostyline III

RL: SPN (Synthetic preparation); PREP (Preparation)
(enantio- and diastereoselective synthesis of substituted tetrahydroisoquinolines)

RN 22325-16-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



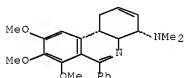
OSC.G 8 THERE ARE 8 CAPLUS RECORDS THAT CITE THIS RECORD (8 CITINGS)

L5 ANSWER 27 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

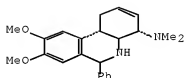
AN 1992:426304 CAPLUS [Full-text](#)

DN 117:26304

OREF 117:4739a,4742a
 TI 1,4,4a,10b-Tetrahydro-N,N-dimethyl-4-phenanthridinamines and
 1,4,4a,5,6,10b-hexahydro-N,N-dimethyl-4-phenanthridinamines
 AU Bobowski, George; West, Barbara; Omecinsky, Diana
 CS Parke-Davis Pharm. Res. Div., Warner-Lambert Co., Ann Arbor, MI, 48105,
 USA
 SO Journal of Heterocyclic Chemistry (1992), 29(1), 33-49
 CODEN: JHTCAD; ISSN: 0022-152X
 DT Journal
 LA English
 OS CASREACT 117:26304
 GI



V



VI

AB Synthetic procedures to prepare the title compds. are described. Diels-Alder cycloaddn. of β -nitrostyrene derivs. to N,N-dimethyl-1,3-butadien-1-amine gave 5-aryl-N,N-dimethyl-6-nitro-2-cyclohexen-1-amines (I). Reduction of I with zinc in acetic acid gave the diamino derivs. II. Schotten-Baumann acylation of II gave amides III. Treatment of II with alkyl isocyanates gave the aminourea derivs. IV. Bischler-Napieralski cyclodehydration of III and IV gave 1,4,4a,10b-tetrahydrophenanthridinamines, e.g., V, and N6-alkyl-1,4,4a,10b-tetrahydro-N4,N4-dimethyl-4,6-phenanthridinediamines, resp. Condensation of diamines II with aryl aldehydes under azeotropic conditions gave imines, which on treatment with acids yielded 6-aryl-1,4,4a,5,6,10b-hexahydro-N,N-dimethyl-4-phenanthridinamines, e.g., VI. The stereochem. of these materials is assigned by NMR.

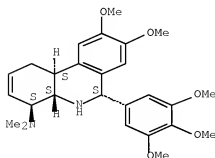
IT 141944-66-3P

RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)

RN 141944-66-3 CAPLUS

CN 4-Phenanthridinamine, 1,4,4a,5,6,10b-hexahydro-8,9-dimethoxy-N,N-dimethyl-6-(3,4,5-trimethoxyphenyl)-, (4a,4aa,6 β ,10b β)- (9CI)
 (CA INDEX NAME)

Relative stereochemistry.



OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)

L5 ANSWER 28 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1992:214753 CAPLUS Full-text

DN 116:214753

OREF 116:36409a,36412a

TI Enantioselective synthesis of 1-substituted 1,2,3,4-tetrahydroisoquinoline alkaloids via asymmetric reduction

AU Cho, Byung Tae; Han, Cheol Kyu

CS Dep. Chem., Hallym Univ., Chunchon, 200-702, S. Korea

SO Bulletin of the Korean Chemical Society (1991), 12(5), 565-9

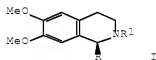
CODEN: BKCSDE; ISSN: 0253-2964

DT Journal

LA English

OS CASREACT 116:214753

GI



AB Enantioselective synthesis of 1-substituted tetrahydroisoquinoline alkaloids I (R = alkyl, aryl, aralkyl; R1 = H, Me) via asym. reduction of 1-substituted 3,4-dihydroisoquinolines and the corresponding iminium salts with the selected chiral hydride reagents, such as K glucuronide Itsuno's reagent and Mosher's reagent were examined. In these reactions, dihydroisoquinolines were not reduced by the hydride reagents, whereas the iminium salts were easily reduced under the same reaction conditions found in successful reduction of ketones. Thus, the reduction of 6,7-dimethoxy-3,4-dihydroisoquinolinium iodide with the chiral reducing agents provided the product I (R = R1 = Me) with 52.3% ee, 18% ee, and 66.4% ee, resp.

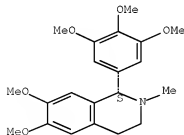
IT 22325-16-2P 33033-86-2P

RL: RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)
(enantioselective synthesis of, via asym. reduction)

RN 22325-16-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

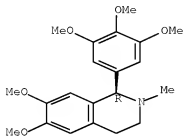


RN 33033-86-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-

trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)

L5 ANSWER 29 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1992:151593 CAPLUS [Full-text](#)

DN 116:151593

OREF 116:25653a,25656a

TI Preparation of tetrahydroisoquinolines as intermediates for alkaloids

IN Takano, Seiichi; Suzuki, Masato; Ogasawara, Kuniro

PA Kawaken Fine Chemicals Co., Ltd., Japan

SO Jpn. Kokai Tokkyo Koho, 3 pp.

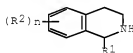
CODEN: JKXXAF

DT Patent

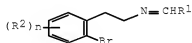
LA Japanese

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|----------------------------------------|------|----------|-----------------|--------------|
| PI | JP 03236375 | A | 19911022 | JP 1990-30698 | 19900209 <-- |
| PRAI | JP 1990-30698 | | 19900209 | | |
| OS | CASREACT 116:151593; MARPAT 116:151593 | | | | |
| GI | | | | | |



I



II

AB The title compds. I (R1 = aryl; R2 = H, alkyl, alkoxy; n = 1, 2), are prepared by treating aromatic imines II (R1, R2, n = same as I) with radical initiators and Bu3SnH in solvents. II [R1 = Ph, (R2)n = 4,5-dimethoxy], Bu3SnH, and AIBN in toluene were refluxed for 2 h to give 21.1% I [R1 = Ph, (R2)n = 6,7-dimethoxy].

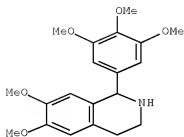
IT 33033-84-0P

RL: SPN (Synthetic preparation); PREP (Preparation)

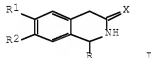
(preparation of, by cyclization of benzyldiene(bromophenyl)ethylamine)

RN 33033-84-0 CAPLUS

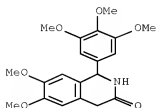
CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
(CA INDEX NAME)



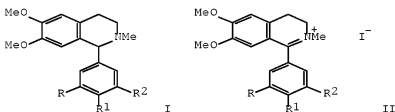
L5 ANSWER 30 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1990:591122 CAPLUS [Full-text](#)
 DN 113:191122
 OREF 113:32349a,32352a
 TI Substituted 1,4-dihydro-1-phenylisoquinolin-3(2H)-ones as inhibitors of
 cyclic nucleotide phosphodiesterases from dog heart
 AU Georgiev, V. S.; Van Inwegen, R. G.; Carlson, P.
 CS Rorer Cent. Res., Horsham, PA, 19044, USA
 SO European Journal of Medicinal Chemistry (1990), 25(4), 375-8
 CODEN: EJMCA5; ISSN: 0223-5234
 DT Journal
 LA English
 OS CASREACT 113:191122
 GI



AB Phenylisoquinolinone and -thione derivs. I [R = Ph, 3,4,5-(MeO)3C6H2; R1, R2 =
 H, MeO; X = O, S] were prepared and tested for their antiallergic activity.
 1-Phenylisoquinoline derivs. also showed antiallergic activity.
 IT 130042-78-3P
 RL BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use);
 BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation and antiallergic activity of)
 RN 130042-78-3 CAPLUS
 CN 3(2H)-Isoquinolinone, 1,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
 (CA INDEX NAME)

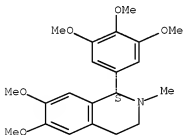


L5 ANSWER 31 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1990:497865 CAPLUS [Full-text](#)
 DN 113:97865
 OREF 113:16545a,16548a
 TI Enantioselective synthesis of cryptostyline I, II and III via asymmetric reduction
 AU Cho, Byung Tae; Han, Cheol Kyu
 CS Dep. Chem., Hallym Univ., Chuncheon, 200-702, S. Korea
 SO Bulletin of the Korean Chemical Society (1990), 11(1), 81-2
 CODEN: BKCSDE; ISSN: 0253-2964
 DT Journal
 LA English
 GI



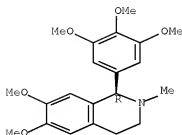
AB The title compds. [(S)-I, R = H, R1R2 = OCH2O, R1 = R2 = MeO; R = R1 = R2 = MeO] were prepared by asym. reduction of the isoquinolinium salts II with K glucoride, Itsuno's reagent, and Maskerh's reagent.
 IT 22325-16-2P, Cryptostyline III 33033-86-2P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)
 RN 22325-16-2 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



RN 33033-86-2 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)

L5 ANSWER 32 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1990:497864 CAPLUS Full-text

DN 113:97864

OREF 113:16545a,16548a

TI Synthesis of racemic cryptostylinines I, II, and III by radical cyclization

AU Takano, Seiichi; Suzuki, Mahito; Kijima, Atsushi; Ogasawara, Kunio

CS Pharm. Inst., Tohoku Univ., Sendai, 980, Japan

SO Chemistry Letters (1990), (2), 315-16

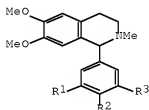
CODEN: CMLTAG; ISSN: 0366-7022

DT Journal

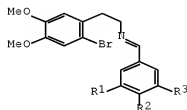
LA English

OS CASREACT 113:97864

GI



I



II

AB Three 1-phenyl-1,2,3,4-tetrahydroisoquinoline alkaloids, cryptostylinines I (I, R1R2 = OCH2O, R3 = H), II (I, R1 = R2 = MeO, R3 = H), and III (I, R1 = R2 = R3 = MeO) were synthesized in racemic forms via a aryl radical-initiated cyclization of imines II.

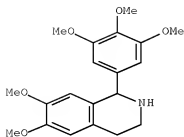
IT 33033-84-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

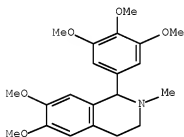
(preparation and methylation of)

RN 33033-84-0 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)

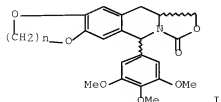


IT 22324-83-0F, (±)-Cryptostyline III
 RL: RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)
 (synthesis of)
 RN 22324-83-0 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



OSC.G 28 THERE ARE 28 CAPLUS RECORDS THAT CITE THIS RECORD (28 CITINGS)

L5 ANSWER 33 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1990:458769 CAPLUS Full-text
 DN 113:58769
 OREF 113:9931a,9934a
 TI The synthesis of 4-desoxy-2-azapodophyllotoxins
 AU Van der Eycken, J.; Bosmans, J. P.; Van Haver, D.; Vandewalle, M.;
 Hulkenberg, A.; Veerman, W.; Nieuwenhuizen, R.
 CS Dep. Org. Chem., State Univ. Gent, Ghent, B-9000, Belg.
 SO Tetrahedron Letters (1989), 30(29), 3873-6
 CODEN: TELEAY; ISSN: 0040-4039
 DT Journal
 LA English
 OS CASREACT 113:58769
 GI



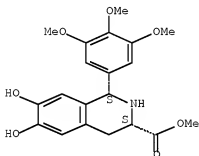
AB 4-Desoxy-2-azapodophyllotoxins I (n = 1, 2), tetrahydroisoquinoline analogs of podophyllotoxin, have been synthesized and evaluated for their antitumor activities.

IT 123049-96-7P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation and reaction of, with dibromoethane)

RN 128049-96-7 CAPLUS

CN 3-Isoquinolinecarboxylic acid, 1,2,3,4-tetrahydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, methyl ester, (1S-cis)- (9CI) (CA INDEX NAME)

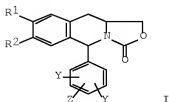
Absolute stereochemistry.



OSC.G 16 THERE ARE 16 CAPLUS RECORDS THAT CITE THIS RECORD (16 CITINGS)

L5 ANSWER 34 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1990:118803 CAPLUS Full-text
 DN 112:118803
 OREF 112:20131a,20134a
 TI Preparation of oxazoloisoquinolines as anticancer agents
 IN Tawara, Tetsuji; Ichanagi, Yukio; Yamagami, Keiji; Fujii, Akihiro
 PA Yoshitomi Pharmaceutical Industries, Ltd., Japan
 SO Jpn. Kokai Tokkyo Koho, 9 pp.
 CODEN: JKXXAF
 DT Patent
 LA Japanese
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|-------------------|------|----------|-----------------|--------------|
| PI | JP 01199976 | A | 19890811 | JP 1987-284890 | 19871110 <-- |
| PRAI | JP 1987-255755 | A1 | 19871008 | | |
| OS | MARPAT 112:118803 | | | | |
| GI | | | | | |



AB The title compds. I (R1, R2 = OH, alkoxy, R1R2 may be alkylenedioxy; X, Y, Z = OH, alkoxy), which have low toxicity, are prepared Treatment of 15.0 g Me cis-2-benzoyloxycarbonyl-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)- 1,2,3,4-tetrahydroisoquinoline-3-carboxylate (preparation given) with CH2Br2, K2CO3, and CuO in DMF at 100-110° for 4 h gave 12.0 g Me cis-2-benzoyloxycarbonyl-1-(3,4,5-trimethoxyphenyl)-6,7-methylenedioxy- 1,2,3,4-tetrahydroisoquinoline-3-carboxylate, which (3.4 g) was treated with LiBH4 in THF at 50-55° for 9 h to afford 1.8 g cis-I (R1R2 = OCH2O; X = 3-OMe, Y = 4-OMe, Z = 5-OMe) (II). Administration of II at 100 mg/kg/day i.p. for 5 days to P388 leukemia cell-bearing mice showed 210% T/C (treated group/control group) survival time, vs. 157%, for podophylotoxin at 25 mg/kg/day. II at 300 mg/kg (i.p.) or 1000 mg/kg (p.o.) exhibited no toxicity (no information on test animal), whereas podophylotoxin at 25 mg/kg was toxic.

IT 125445-07-0P 125445-08-1F 125445-09-2F
125445-10-5P 125445-13-6F 125445-14-9F
125445-15-0P 125456-07-7P

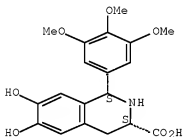
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation and reaction of, in preparation of oxazoloisoquinolines)

RN 125445-07-0 CAPLUS

CN 3-Isoquinolinecarboxylic acid, 1,2,3,4-tetrahydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, cis- (9CI) (CA INDEX NAME)

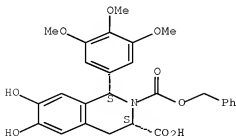
Relative stereochemistry.



RN 125445-08-1 CAPLUS

CN 2,3(1H)-Isoquinolinedicarboxylic acid, 3,4-dihydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, 2-(phenylmethyl) ester, cis- (9CI) (CA INDEX NAME)

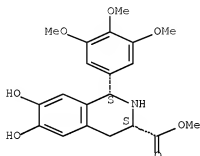
Relative stereochemistry.



RN 125445-09-2 CAPLUS

CN 3-Isoquinolinecarboxylic acid, 1,2,3,4-tetrahydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, methyl ester, hydrochloride, cis- (9CI) (CA INDEX NAME)

Relative stereochemistry.

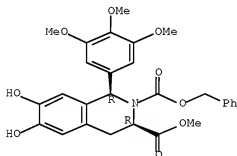


● HCl

RN 125445-10-5 CAPLUS

CN 2,3(1H)-Isoquinolinedicarboxylic acid, 3,4-dihydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, 3-methyl 2-(phenylmethyl) ester, cis- (9CI) (CA INDEX NAME)

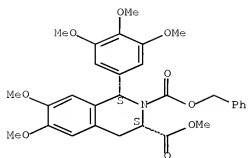
Relative stereochemistry.



RN 125445-13-8 CAPLUS

CN 2,3(1H)-Isoquinolinedicarboxylic acid, 3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, 3-methyl 2-(phenylmethyl) ester, cis- (9CI) (CA INDEX NAME)

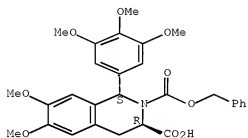
Relative stereochemistry.



RN 125445-14-9 CAPLUS

CN 2,3(1H)-Isoquinolinedicarboxylic acid,
3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, 2-(phenylmethyl)
ester, trans- (9CI) (CA INDEX NAME)

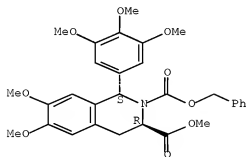
Relative stereochemistry.



RN 125445-15-0 CAPLUS

CN 2,3(1H)-Isoquinolinedicarboxylic acid,
3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, 3-methyl
2-(phenylmethyl) ester, trans- (9CI) (CA INDEX NAME)

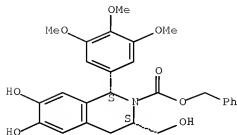
Relative stereochemistry.



RN 125456-07-7 CAPLUS

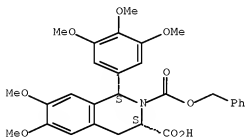
CN 2(1H)-Isoquinolinecarboxylic acid,
3,4-dihydro-6,7-dihydroxy-3-(hydroxymethyl)-1-(3,4,5-trimethoxyphenyl)-,
phenylmethyl ester, cis- (9CI) (CA INDEX NAME)

Relative stereochemistry.



IT 125456-06-8P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)
 RN 125456-08-8 CAPLUS
 CN 2,3(1H)-Isoquinolinedicarboxylic acid,
 3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, 2-(phenylmethyl)
 ester, cis- (9CI) (CA INDEX NAME)

Relative stereochemistry.



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

L5 ANSWER 35 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1989:497608 CAPLUS [Full-text](#)

DN 111:97608

OREF 111:16437a,16440a

TI Carbon-transfer reactions with heterocycles. Part 3. Synthetic
 equivalence of oxazolidines with carbonyl compounds

AU Singh, Harjit; Sarin, Rakesh

CS Dep. Chem., Guru Nanak Dev Univ., Amritsar, 143005, India

SO Journal of Chemical Research, Synopses (1986), (10), 322-3

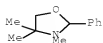
CODEN: JRPSDC; ISSN: 0308-2342

DT Journal

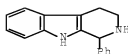
LA English

OS CASREACT 111:97608

GI



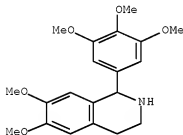
I



II

AB Oxazolidinones transfer C-2 carbon units to binucleophiles to furnish heterocycles including β -carboline and isoquinoline alkaloids. Thus, the oxazolidine I was treated with tryptamine in MeCN containing acid to give its carboline II.

IT 33033-84-0P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)
 RN 33033-84-0 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
 (CA INDEX NAME)



OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)

L5 ANSWER 36 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1989:154166 CAPLUS Full-text

DN 110:154166

OREF 110:25491a,25494a

TI Preparation of di- and tetrahydroisoquinoline derivatives as cytostatics

PA Duphar International Research B. V., Neth.

SO Jpn. Kokai Tokkyo Koho, 13 pp.

CODEN: JKXXAF

DT Patent

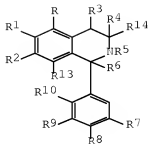
LA Japanese

FAN.CNT 1

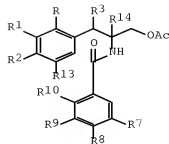
| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|-------------------------------------------------------|------|----------|-----------------|--------------|
| PI | JP 62283960 | A | 19871209 | JP 1987-119169 | 19870518 <-- |
| | CA 1330560 | C | 19940705 | CA 1987-537195 | 19870515 <-- |
| | DK 8702503 | A | 19871122 | DK 1987-2503 | 19870518 <-- |
| | AU 8773149 | A | 19871126 | AU 1987-73149 | 19870518 <-- |
| | ZA 8703561 | A | 19871230 | ZA 1987-3561 | 19870518 <-- |
| | IL 82555 | A | 19910512 | IL 1987-82555 | 19870518 <-- |
| | EP 251361 | A1 | 19880107 | EP 1987-200926 | 19870519 <-- |
| | R: AT, BE, CH, DE, ES, FR, GB, GR, IT, LI, LU, NL, SE | | | | |
| | US 5162335 | A | 19921110 | US 1989-480755 | 19891127 <-- |
| PRAI | NL 1986-1279 | A | 19860521 | | |
| | US 1987-50425 | B2 | 19870518 | | |
| | US 1987-123326 | B1 | 19871120 | | |

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

GI



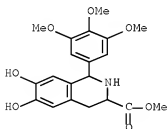
I



II

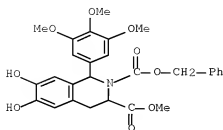
AB Title compds. I [R, R1, R2 = H, halo, CF₃, C1-4 alkyl, C1-4 alkoxy; R1R1, R1R2, R2R13 = hydrocarbenyl (may be interrupted with 1-3 N, O, or S) to form 5-7 membered cycle which may be substituted with C1-4 alkyl, halo, oxo, or thioxo; R3 = H, OH, C1-4 alkoxy, alkanoyloxy; R4 = H, CO₂R11, CON(R12)2, CH₂OR15 (R11, R12 = C1-4 alkyl; R15 = H, C1-4 alkyl, tetrahydropyranyl, C1-4 alkanoyl, mono- or di-C1-4 alkyl substituted carbamoyl); R5 = H, C1-4 alkanoyl, alkyl; R4R5 = CH₂XZ (X = O, NH, alkyl-substituted N; Z = CO, CS, SO, CH₂); R6 = H; R5R6 = bond; R7, R9 = H, C1-4 alkoxy; R8 = H, C1-4 alkoxy, alkanoyloxy, OH, halo; R7R8, R8R9 = OCH₂O, O(CH₂)₂O; R10 = H, C1-4 alkyl, alkoxy; R14 = H, Me] are prepared from N-phenethylbenzamides II. A solution of II (R = R3 = R10 = R13 = R14 = H; R1R2 = OCH₂O; R7 = R8 = R9 = MeO) was successively treated with PC15 and AlCl₃ to give 92% I (R = R3 = R10 = R13 = R14 = H; R1R2 = OCH₂O; R4 = AcOCH₂; R5R6 = bond; R7 = R8 = R9 = MeO), which was hydrolyzed with K₂CO₃ in MeOH to afford 83% I (R4 = CH₂OH). The latter product in THF was treated with AlH₃ (generated from AlCl₃ and LiAlH₄ in situ) for 1.5 h to give 97% I (R5 = R6 = H), which in CH₂Cl₂ was treated with Cl₂CO in the presence of NEt₃ to afford 89% I [R = R3 = R6 = R10 = R13 = R14 = H; R1R2 = OCH₂O; R4R5 = CH₂OC(O); R7 = R8 = R9 = MeO; cis-form at 1,3-position] (III). III at 15 μg/mL showed .apprx.50-100% control of human cell lines.

IT 118068-19-2P 118068-20-5P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation and reaction of, in preparation of isoquinoline cytostatics)
 RN 118068-19-2 CAPLUS
 CN 3-Isoquinolinecarboxylic acid, 1,2,3,4-tetrahydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, methyl ester (CA INDEX NAME)



RN 118068-20-5 CAPLUS
 CN 2,3(1H)-Isoquinolinedicarboxylic acid,
 3,4-dihydro-6,7-dihydroxy-1-(3,4,5-trimethoxyphenyl)-, 3-methyl

2-(phenylmethyl) ester (CA INDEX NAME)



OSC.G 6 THERE ARE 6 CAPLUS RECORDS THAT CITE THIS RECORD (6 CITINGS)

L5 ANSWER 37 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1984:610940 CAPLUS [Full-text](#)

DN 101:210940

OREF 101:31959a

TI Synthesis and radioprotectant properties of some derivatives of 6-hydroxy-1,2,3,4-tetrahydroisoquinoline

AU Alpatova, T. V.; Klimova, A. D.; Kulinskii, V. I.; Mirzoyan, V. S.; Mirzoyan, A. T.; Yashunskii, V. G.

CS Inst. Biofiz., Moscow, USSR

SO Khimiko-Farmatsevticheskii Zhurnal (1984), 18(4), 444-9

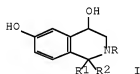
CODEN: KHFZAN; ISSN: 0023-1134

DT Journal

LA Russian

OS CASREACT 101:210940

GI



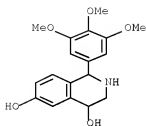
AB Cyclocondensation of 3-HOC6H4CH(OH)CH2NHCH2Ph with R2CHO (R2 = Me, 2-, 4-HOC6H4, 4-pyridyl) gave 36.5-88% isoquinolinols I (R = PhCH2, R1 = H, R2 as above) which were debenzylated by Pd-C to give 47-76% I (R = R1 = H, R2 = Me, 2-, 4-HOC6H4). Cyclocondensation of 3-HOC6H4CH(OH)CH2NHMe with R2CHO gave 30-50% I (R = Me, R1 = H, R2 = H, 2-HOC6H4, 4-MeOC6H4). Addnl. obtained were 3-HOC6H4CH(OH)CH2NRCHR1R2 (R = R1 = H, R2 = Me, 4-MeOC6H4; R = Me, R1 = R2 = H; R = H, R1 = R2 = Me). Mice treated with I (R = PhCH2, R1 = H, R2 = Me) (LD50 3521 ± 56 μmol/kg) increased their survival time to 13 ± 6% at 50 μmol/kg dosage compared to 5.6 ± 1.3 for a control.

IT 93202-93-9 93202-96-1

RL: RCT (Reactant); RACT (Reactant or reagent)
(radioprotectant properties of)

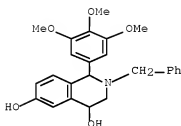
RN 93202-94-9 CAPLUS

CN 4,6-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



RN 93202-96-1 CAPLUS

CN 4,6-Isoquinolinediol, 1,2,3,4-tetrahydro-2-(phenylmethyl)-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

L5 ANSWER 38 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1984:530578 CAPLUS [Full-text](#)

DN 101:130578

OREF 101:19861a,19864a

TI Synthesis of benzazepinespirocycloalkanes. IV.

1,2-Substituted-1,2,3,4-tetrahydrospiro-5-cyclopentane-(5H)-2-benzazepines

AU Solomina, L. P.; Pirdzhanov, L. Sh.; Markaryan, E. A.

CS Inst. Tonkoi Org. Khim. im. Mndzhoyana, Yerevan, USSR

SO Armyanskii Khimicheskii Zhurnal (1984), 37(4), 253-7

CODEN: AYKZAN; ISSN: 0515-9628

DT Journal

LA Russian

OS CASREACT 101:130578

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Condensation of cyclopentaneethanamine I and spiro[benzazepine-cyclopentane] II with 3,4,5-(MeO)2RC6H2(CH2)nCOCl (R = MeO, n = 0; R = H, n = 1) yields amides III and IV. Cyclodehydration and reduction of III with POC13 and NaBH4 gives V [R1 = 3,4-(MeO)2C6H3CH2, 3,4,5-(MeO)3C6H2, R2 = H] which were N-methylated by CH2O-HCO2H to give V (R2 = Me). Reduction of IV by LiAlH4 gave 50.0 and 42.3% V [R1 = Me, R2 = 3,4-(MeO)2C6H3CH2CH2, 3,4,5-(MeO)3C6H2CH2].

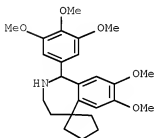
IT 91935-74-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation and N-methylation of)

RN 91935-74-9 CAPLUS

CN Spiro[5H-2-benzazepine-5,1'-cyclopentane],
1,2,3,4-tetrahydro-7,8-dimethoxy-1-(3,4,5-trimethoxyphenyl)-,
hydrochloride (1:1) (CA INDEX NAME)



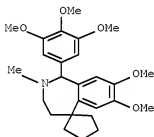
● HCl

IT 91935-76-1P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 91935-76-1 CAPLUS

CN Spiro[5H-2-benzazepine-5,1'-cyclopentane],
1,2,3,4-tetrahydro-7,8-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-,
hydrochloride (1:1) (CA INDEX NAME)



● HCl

OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

L5 ANSWER 39 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1982:509845 CAPLUS [Full-text](#)

DN 97:109845

OREF 97:18269a,18272a

TI A new variant of the internal α -amidoalkylation reaction: synthesis
of 1-aryl-3-oxo-2,3-dihydroisoquinolines and cryptostyline III

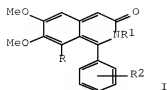
AU Venkov, A.; Lukanov, L.; Mollov, N.

CS Dep. Chem., Univ. Plovdiv, Plovdiv, 4000, Bulg.

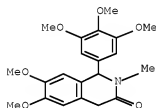
SO Synthesis (1982), (6), 486-7

CODEN: SYNTBF; ISSN: 0039-7881

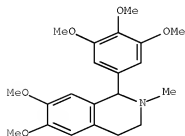
DT Journal
LA English
GI



- AB Isoquinolinones I [R = H, OMe; R1 = Me, Ph, 4-MeOC6H4; R2 = H, 4-O2N, 4-Cl, 4-F, 3,4,5-(MeO)3] were obtained at 32-67% yield by treating 3,4,5-R(MeO)2C6H2CH2CO2H with R2C6H4CONHR1 in the presence of POCl3. Cryptostyline III was prepared by 2-stage reduction of I [R = H, R1 = Me, R2 = 3,4,5-(MeO)3].
- IT 92801-29-3F
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation and reduction of)
- RN 82801-28-3 CAPLUS
- CN 3(2H)-Isoquinolinone, 1,4-dihydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)

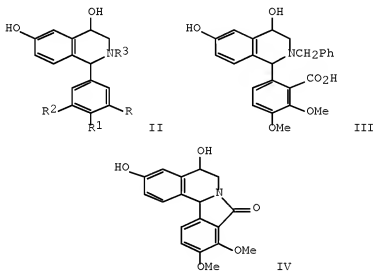


- IT 22324-83-0F
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)
- RN 22324-83-0 CAPLUS
- CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



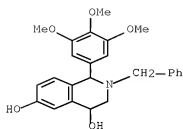
OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)

L5 ANSWER 40 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1981:603702 CAPLUS Full-text
 DN 95:203702
 OREF 95:34033a
 TI Use of a benzyl protective group in the synthesis of
 tetrahydroisoquinoline derivatives
 AU Alpatova, T. V.; Yashunskii, V. G.
 CS Inst. Biofiz., Moscow, 123182, USSR
 SO Khimiya Geterotsiklicheskih Soedinenii (1981), (8), 1084-7
 CODEN: KGSSAQ; ISSN: 0453-8234
 DT Journal
 LA Russian
 OS CASREACT 95:203702
 GI



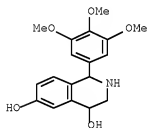
AB Cyclocondensation of 3-HOC6H4CH(OH)CH2NHCH2Ph (I) with 3,4,5-RR1R2C6H2CHO (R,R1,R2 = MeO,H,O; MeO,MeO,MeO; H,Me2N,H) gave benzylisoquinolines II (R3 = benzyl), which were debenzylated by hydrogenolysis over Pd black to give II (R3 = H). I and CH2O gave a mixture of 2-benzyl-1,2,3,4-tetrahydroisoquinoline-4,6-diol and 2-benzyl-1,2,3,4-tetrahydroisoquinoline-4,8-diol; these compds. were also debenzylated. Debenzylation-cyclocondensation of III gave lactam IV.

IT 79677-06-8F
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation and debenzylation of)
 RN 79677-06-8 CAPLUS
 CN 4,6-Isoquinolinediol, 1,2,3,4-tetrahydro-2-(phenylmethyl)-1-(3,4,5-trimethoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME)



● HCl

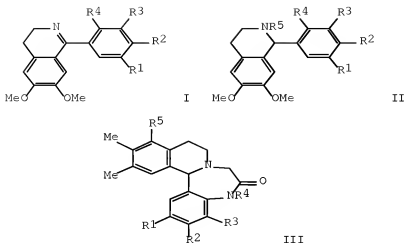
IT 72512-01-7P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)
 RN 72512-01-7 CAPLUS
 CN 4,6-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)-,
 hydrochloride (1:1) (CA INDEX NAME)



● HCl

OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)

L5 ANSWER 41 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1981:198 CAPLUS Full-text
 DN 94:198
 OREF 94:39a,42a
 TI Pharmacological studies on some derivatives of the isoquinoline series
 AU Nikolova, M.; Ivanova, N.
 CS Bulg.
 SO Trudove na Nauchnoizsledovatel'skiya Khimikofarmatsevtichen Institut (1978), 10, 211-19
 CODEN: TKZGAG; ISSN: 0371-8972
 DT Journal
 LA Bulgarian
 GI



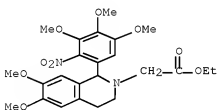
AB Among 12 title compds. I (R1 and R2 and R3 = H, Cl, or MeO; R4 = NO2, NH2, or NHCOCBr(Me)2) II (R1 and R2 and R3 = MeO; R4 = NO2; R5 = CH2CO2Et), and III (R1 and R2 and R3 = H, Cl; R4 = H, Et, CHMe2; R5 = H, NH2, NO2), evaluated pharmacol. in vivo, III showed weak central depressive activity and lower toxicity than I. All compds. had low hypotensive activity and little effect on the smooth muscle of gastrointestinal tract and bronchi.

IT 75230-95-4

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(pharmacol. of)

RN 75230-95-4 CAPLUS

CN 2(1H)-Isoquinolineacetic acid, 3,4-dihydro-6,7-dimethoxy-1-(3,4,5-trimethoxy-2-nitrophenyl)-, ethyl ester (CA INDEX NAME)



L5 ANSWER 42 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1980:531967 CAPLUS Full-text

DN 93:131967

OREF 93:21029a, 21032a

TI Carbon-13 NMR spectra of 4-hydroxytetrahydroisoquinolinium chlorides

AU Baddeley, G. Vernon; Quessy, Stephen N.; Williams, Lyall R.

CS Sch. Chem., Univ. New South Wales, Kensington, 2033, Australia

SO Australian Journal of Chemistry (1980), 33(2), 447-50

CODEN: AJCHAS; ISSN: 0004-9425

DT Journal

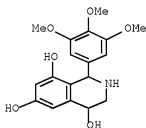
LA English

AB The natural abundance ^{13}C NMR spectra of tetrahydroisoquinolinium salts are reported and discussed in terms of the substitution profile and stereochem. of the salts.

IT 72511-98-9 72512-01-7
 RL: PRP (Properties)
 (carbon-13 NMR of)

RN 72511-98-9 CAPLUS

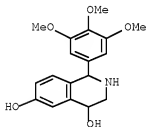
CN 4,6,8-Isoquinolinetriol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME)



● HCl

RN 72512-01-7 CAPLUS

CN 4,6-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME)



● HCl

L5 ANSWER 43 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1980:58575 CAPLUS [Full-text](#)

DN 92:58575

OREF 92:9699a,9702a

TI Synthesis and biological evaluation of tetrahydroisoquinolin-4-ol derivatives

AU Quessy, Stephen N.; Williams, Lyall R.

CS Sch. Chem., Macquarie Univ., North Ryde, 2113, Australia

SO Australian Journal of Chemistry (1979), 32(6), 1317-27

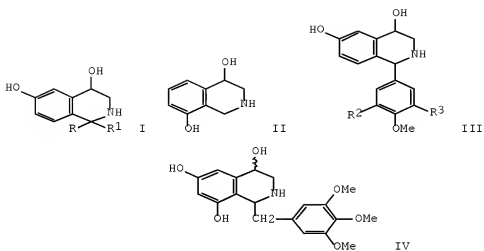
CODEN: AJCHAS; ISSN: 0004-9425

DT Journal

LA English

OS CASREACT 92:58575

GI



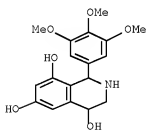
AB Hydroxytetrahydroisoquinolines I (R = R₁ = H, Me), II, III (R₂, R₃ = H, MeO), and IV, cyclic analogs of phenylethanalamines, were prepared I-IV had no β -adrenoceptor stimulating activity.

IT 72511-96-9P 72512-01-7P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 72511-98-9 CAPLUS

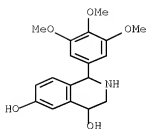
CN 4,6,8-Isoquinolinetriol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME)



● HCl

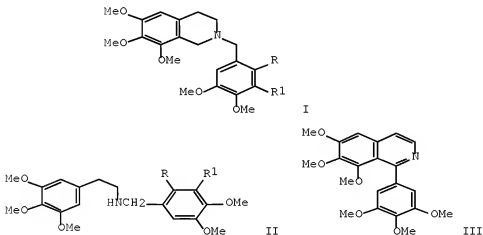
RN 72512-01-7 CAPLUS

CN 4,6-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME)



● HC1

L5 ANSWER 44 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1978:121486 CAPLUS [Full-text](#)
 DN 88:121486
 OREF 88:19081a,19084a
 TI Polymethoxylated isoquinolines as potential antimitotic agents
 AU Torio, Maria; Brossi, Arnold; Chignell, Colin F.
 CS Lab. Chem., Natl. Inst. Arthritis, Metab. Dig. Dis., Bethesda, MD, USA
 SO Heterocycles (1978), 9(1), 1-6
 CODEN: HICYAM; ISSN: 0385-5414
 DT Journal
 LA English
 GI



AB The sendaverine derivs. I (R = MeO, R1 = H; R = H, R1 = MeO) were prepared by condensation of mescaline with 2,4,5-(MeO)3C6H2CHO and 3,4,5-(MeO)3C6H2CHO and reduction of the Schiff bases to give II, which were cyclized. The cryptostyline derivative III was prepared by cyclization of 3,4,5-(MeO)3C6H2CONHCH2CH2C6H2(OMe)3-3,4,5 followed by reduction and dehydrogenation. Using colchicine as a standard none of the compds. showed any binding affinity to the rat brain microtubule protein.

IT 65967-39-7P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation and dehydration of)

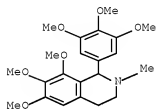
RN 65967-39-7 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7,8-trimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, ethanedioate (1:?) (CA INDEX NAME)

CM 1

CRN 65967-38-6

CMF C22 H29 N O6



CM 2

CRN 144-62-7

CMF C2 H2 O4

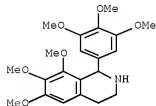


IT 65967-37-5P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)
(preparation and methylation of)

RN 65967-37-5 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7,8-trimethoxy-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



L5 ANSWER 45 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

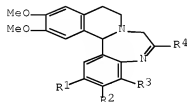
AN 1977:121306 CAPLUS [Full-text](#)

DN 86:121306

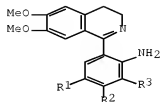
OREF 86:19159a,19162a

TI 1,4-Benzodiazepines. IV. 6-Phenyl- and
6-thienyltetrahydroisoquinolino[2,1-d]-7H-[1,4]-benzodiazepines

AU Ivanov, Ch.; Shvedov, V. I.
 CS Sci.-Res. Chem.-Pharm. Inst., Sofia, Bulg.
 SO Khimiko-Farmatsevticheski Zhurnal (1976), 10(7), 44-51
 CODEN: KHFZAN; ISSN: 0023-1134
 DT Journal
 LA Russian
 GI

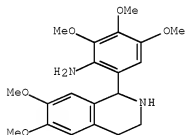


I



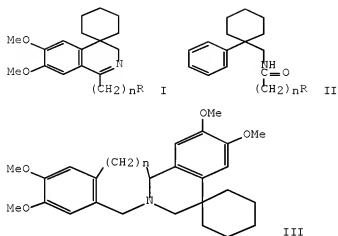
II

AB Isoquinolinobenzodiazepines I (R1 = Cl, H, MeO, R2 = H, Cl, MeO, R3 = H, MeO, R4 = Ph, 4-biphenyl, 4-BrC6H4, 4-MeOC6H4, 4-O2NC6H4, 2-thienyl) were obtained in 15-33% yields by reduction of II with NaBH4 followed by cyclization with R4COCH2Br. Addnl. I (R1 = Cl, R2 = R3 = H, R4 = Ph, 4-biphenyl, 4-BrC6H4, 4-MeOC6H4, 4-O2NC6H4) were obtained from II by cyclization with R4COCH2Br followed by reduction with NaBH4.
 IT 62206-18-2P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation and cyclization by phenacyl bromides)
 RN 62206-18-2 CAPLUS
 CN Benzenamine, 2,3,4-trimethoxy-6-(1,2,3,4-tetrahydro-6,7-dimethoxy-1-isoquinolinyl)- (CA INDEX NAME)



L5 ANSWER 46 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1976:164575 CAPLUS [Full-text](#)
 DN 84:164575
 OREF 84:26711a,26714a
 TI Isoquinoline derivatives. XI. Synthesis and pharmacological activity of 1-arylalkyl-4-spirocyclohexane-6,7-dimethoxy-1,2,3,4-tetrahydroisoquinolines and some of their derivatives
 AU Markaryan, E. A.; Arustamyan, Zh. S.; Vasilyan, S. S.; Markaryan, K. Zh.
 CS Inst. Tonkol Org. Khim. im. Mndzhoyana, Yerevan, USSR
 SO Armysanskii Khimicheskii Zhurnal (1975), 28(10), 829-35
 CODEN: AYKZAN; ISSN: 0515-9628

DT Journal
 LA Russian
 OS CASREACT 84:164575
 GI



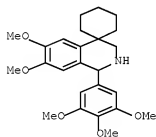
AB Spiro[cyclohexaneisoquinoline] derivs. I [R = Ph, 3,4-(MeO)₂C₆H₃, 3,4,5-(MeO)₃C₆H₂, Ph₂CH, n = 0,1] were obtained in 45.3-69.4% yields by cyclization of amides II with POC13. Reduction of I by NaBH₄ gave 67.3-81.2% 1,2-dihydro derivs. Cyclocondensation of the appropriate 1,2-dihydro derivative with HCHO gave 55.1 and 56% berbines III (n = 1,2). I were useful in treatment of hypertension, as muscle relaxants, and as antiarrhythmics.

IT 59021-77-1F

RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)

RN 59021-77-1 CAPLUS

CN Spiro[cyclohexane-1,4'-(1'H)-isoquinoline],
 2',3'-dihydro-6',7'-dimethoxy-1'-(3,4,5-trimethoxyphenyl)-, hydrochloride
 (1:1) (CA INDEX NAME)



● HCl

OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)

L5 ANSWER 47 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1976:4818 CAPLUS Full-text

DN 84:4818

OREF 84:813a,816a

TI 6,7-Dihydroxy-1-(3,4,5-trimethoxybenzyl)-1,2,3,4-tetrahydroisoquinoline

IN Vasvari, Mrs. Arpad; Meszaros, Zoltan; Nagy, Gabor; Hermecz, Istvan;

Horvath, Agnes; David, Agoston; Mandi, Attila; Pajor, Aniko

PA Chinoin Gyogyszer es Vegyeszeti Termekek Gyara Rt., Hung.

SO Hung. Teljes, 13 pp.

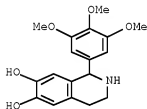
CODEN: HUXXBU

DT Patent

LA Hungarian

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|----------|-----------------|--------------|
| PI | HU 9519 | | 19750328 | HU 1973-CI1389 | 19730628 <-- |
| AB | 3,4-(HO)2C6H3CH2CH2NH2.HBr added to excess Na 3,4,5-trimethoxyphenylglycidate in aqueous HCl-AcOH, the mixture stirred 2 hr at 20° and 5 hr at 70-80° and pH 1 gave 54% title product HCl salt, m. 225-6° (MeOH). | | | | |
| IT | 57529-51-8P RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of) | | | | |
| RN | 57529-51-8 CAPLUS | | | | |
| CN | 6,7-Isoquinolinediol, 1,2,3,4-tetrahydro-1-(3,4,5-trimethoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME) | | | | |



● HCl

L5 ANSWER 48 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1975:593279 CAPLUS Full-text

DN 83:193279

OREF 83:30405a,30408a

TI Heterocyclic compounds

IN Ott, Hans; Suess, Rudolf

PA Sandoz-Patent-G.m.b.H., Fed. Rep. Ger.

SO Ger. Offen., 29 pp.

CODEN: GWXXBX

DT Patent

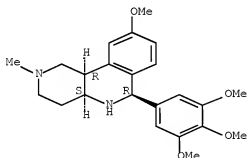
LA German

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|------------|------|----------|-----------------|--------------|
| PI | DE 2503156 | A1 | 19750807 | DE 1975-2503156 | 19750127 <-- |
| | FI 7500205 | A | 19750806 | FI 1975-205 | 19750127 <-- |
| | SE 7500859 | A | 19750806 | SE 1975-859 | 19750127 <-- |
| | DK 7500242 | A | 19750929 | DK 1975-242 | 19750127 <-- |
| | DK 136189 | B | 19770829 | | |

| | | | | |
|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----|----------|----------------|--------------|
| DK 136189 | C | 19780130 | | |
| NO 7500263 | A | 19750806 | NO 1975-263 | 19750129 <-- |
| NL 7501159 | A | 19750807 | NL 1975-1159 | 19750131 <-- |
| BE 825124 | A1 | 19750804 | BE 1975-153007 | 19750203 <-- |
| DD 117216 | A5 | 19760105 | DD 1975-183969 | 19750203 <-- |
| AU 7577843 | A | 19760805 | AU 1975-77843 | 19750203 <-- |
| HU 169923 | B | 19770228 | HU 1975-SA2741 | 19750203 <-- |
| GB 1497722 | A | 19780112 | GB 1975-4516 | 19750203 <-- |
| GB 1497723 | A | 19780112 | GB 1977-30857 | 19750203 <-- |
| JP 50108297 | A | 19750826 | JP 1975-14021 | 19750204 <-- |
| AT 7500799 | A | 19790115 | AT 1975-799 | 19750204 <-- |
| AT 351536 | B | 19790725 | | |
| CA 1055944 | A1 | 19790605 | CA 1975-219303 | 19750204 <-- |
| FR 2259611 | A1 | 19750829 | FR 1975-3547 | 19750205 <-- |
| FR 2259611 | B1 | 19800111 | | |
| ZA 7500743 | A | 19760929 | ZA 1975-743 | 19750205 <-- |
| US 4087530 | A | 19780502 | US 1976-670111 | 19760325 <-- |
| PRAI CH 1974-1550 | A | 19740205 | | |
| US 1975-545540 | A2 | 19750130 | | |
| GB 1975-4516 | A | 19750203 | | |
| GB 1975-18178 | A | 19750917 | | |
| GB 1975-38177 | A | 19750917 | | |
| OS MARPAT 83:193279 | | | | |
| GI For diagram(s), see printed CA Issue. | | | | |
| AB Broncholytic (no data) cis-octahydrobenzonaphthyridines I (R = H, 8-OH, 8-OMe, 9-OMe, 9-OH; R1 = H, 3-OMe, 3-Cl, 3-OH, 2-OH; R2 = H, NH2, NHAc, F, Me, Cl, OMe, OH, NO2, NMe2; R2 = H, OMe, OH) (39 compds.) were prepared by reducing the 1,2,3,4,4a,10b-hexahydro analogs with PtO, or NaBH4. | | | | |
| IT 57251-31-7P | | | | |
| RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of) | | | | |
| RN 57251-31-7 CAPLUS | | | | |
| CN Benzo[c][1,6]naphthyridine, 1,2,3,4,4a,5,6,10b-octahydro-9-methoxy-2-methyl-6-(3,4,5-trimethoxyphenyl)-, dihydrochloride, (4aa,6β,10ba)- (9CI) (CA INDEX NAME) | | | | |

Relative stereochemistry.

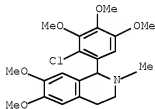


● 2 HCl

OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)

L5 ANSWER 49 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1975:80346 CAPLUS [Full-text](#)
 DN 82:80346

OREF 82:12793a,12796a
 TI Experimental studies on the antitumor effect of ethidium bromide and related substances
 AU Nishiwaki, Hiroshi; Miura, Moriji; Imai, Kuniyuki; Ohno, Ryuzo; Kawashima, Kohei; Ezaki, Koji; Ueda, Ryuzo; Yoshikawa, Haruya; Nagata, Kouichiro; et al.
 CS Sch. Med., Nagoya Univ., Nagoya, Japan
 SO Cancer Research (1974), 34(10), 2699-703
 CODEN: CNREA8; ISSN: 0008-5472
 DT Journal
 LA English
 GI For diagram(s), see printed CA Issue.
 AB Ethidium bromide [1239-45-8] increased by $\leq 200\%$ the life span of mice with tumor 6C3HED-OG, and by $\leq 83\%$ the life span of mice with L5178Y. No such effect was noted in mice with tumors L1210, EL4, 6C3HED-RG, RADA1, or Walker carcinosarcoma 256. Of the 14 related compds. tested, PD-MY-001 (I) [53409-06-6] and PD-MY-003 (II) [38483-26-0] also increased the survival time of mice with 6C3HED-OG by $\leq 200\%$. Ethidium bromide and these 2 newly synthesized compds. are apparently antitumor agents with a unique mechanism of action.
 IT 53954-69-1
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (neoplasm inhibition by)
 RN 53954-69-1 CAPLUS
 CN Isoquinoline, 1-(2-chloro-3,4,5-trimethoxyphenyl)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

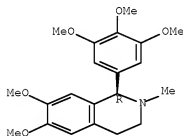


● HCl

OSC.G 5 THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)
 L5 ANSWER 50 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1974:435571 CAPLUS Full-text
 DN 81:35571
 OREF 81:5709a,5712a
 TI Orchidaceae alkaloids. XXXIX. Isolation of (-)-cryptostyline I, II, III and two quaternary salts from *Cryptostylis erythroglossa*. Biosynthetic studies of (-)-cryptostyline I
 AU Agurell, Stig; Granelli, Ingrid; Leander, Kurt; Luning, Bjorn; Rosenblom, Jan
 CS Fac. Pharm., Stockholm, Swed.
 SO Acta Chemica Scandinavica, Series B: Organic Chemistry and Biochemistry (1974), 28(2), 239-43
 CODEN: ACBOCV; ISSN: 0302-4369
 DT Journal
 LA English

- AB (-)-Cryptostyline I, II, and III, together with 1-(3,4-methylenedioxyphenyl)-6,7-dimethoxy-2-methyl-3,4-dihydroisoquinolinium iodide and 1-(3,4-methylenedioxyphenyl)-6,7-dimethoxy-2-methylisoquinolinium chloride have been isolated from *Cryptostylis erythroglossa*. The biosynthesis of (-)-cryptostyline I has been studied using radioactive precursors and the position of the radio-label determined by degradation. The biosynthetic results show that tyrosine and 3,4-dihydroxyphenylalanine as well as tyramine and dopamine are specifically incorporated. The finding that 3-hydroxy-4-methoxyphenethylamine is better incorporated than the isomeric 4-hydroxy-3-methoxy-phenethylamine suggests that the ring closure to the tetrahydroisoquinoline skeleton is facilitated by a para-hydroxy group.
- IT 33033-66-2
 RL: BOC (Biological occurrence); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence)
 (of *Cryptostylis erythroglossa*)
- RN 33033-86-2 CAPLUS
- CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

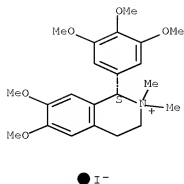
Absolute stereochemistry.



OSC.G 4 THERE ARE 4 CAPLUS RECORDS THAT CITE THIS RECORD (4 CITINGS)

- L5 ANSWER 51 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
- AN 1973:525682 CAPLUS [Full-text](#)
- DN 79:125682
- OREF 79:20407a,20410a
- TI Syntheses of heterocyclic compounds. DXXVII. Absolute configuration of 1,2,3,4-tetrahydro-1-phenylisoquinolines
- AU Kametani, Tetsuji; Shibuya, Shiroshi; Sugi, Hideo; Fukumoto, Keiichiro
- CS Pharm. Inst., Tohoku Univ., Sendai, Japan
- SO Journal of Heterocyclic Chemistry (1973), 10(4), 451-3
 CODEN: JHTCAD; ISSN: 0022-152X
- DT Journal
- LA English
- AB The absolute configuration at the C-1 position of 1,2,3,4-tetrahydro-1-phenylisoquinolines can be deduced from the CD curves of either the free base or its methiodide. The absolute configuration of (+)-2-amino-1-(3-hydroxyphenyl)ethanol was revised and found to have R-configuration.
- IT 43090-64-8
 RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
 (preparation and circular dichroism of)
- RN 43090-64-8 CAPLUS
- CN Isoquinolinium, 1,2,3,4-tetrahydro-6,7-dimethoxy-2,2-dimethyl-1-(3,4,5-trimethoxyphenyl)-, iodide, (S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



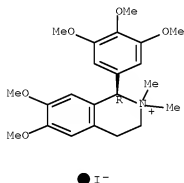
IT 43090-65-9P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 43090-65-9 CAPLUS

CN Isoquinolinium, 1,2,3,4-tetrahydro-6,7-dimethoxy-2,2-dimethyl-1-(3,4,5-trimethoxyphenyl)-, iodide, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



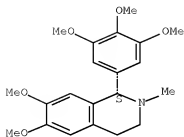
IT 22325-16-2 33033-06-2

RL: RCT (Reactant); RACT (Reactant or reagent)
(quaternization of)

RN 22325-16-2 CAPLUS

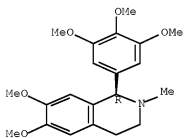
CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



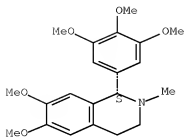
RN 33033-86-2 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L5 ANSWER 52 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1973:159937 CAPLUS [Full-text](#)
 DN 78:159937
 OREF 78:25687a,25690a
 TI Orchidaceae alkaloids. XXXIV. Absolute configuration of cryptostyline I, II, and III. Three 1-phenyl-1,2,3,4-tetrahydroisoquinolines from *Cryptostylyis fulva*
 AU Leander, Kurt; Luning, Bjorn; Westin, Leif
 CS Dep. Org. Chem., Univ. Stockholm, Stockholm, Swed.
 SO Acta Chemica Scandinavica (1947-1973) (1973), 27(2), 710
 CODEN: ACSAA4; ISSN: 0001-5393
 DT Journal
 LA English
 GI For diagram(s), see printed CA Issue.
 AB Cryptostyline I, II and III isolated from *Cryptostylyis fulva* had structures I, II, and III, resp., based on CD and x-ray crystallog.
 IT 22325-16-2
 RL: PRP (Properties)
 (structure and configuration of)
 RN 22325-16-2 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



L5 ANSWER 53 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1973:111557 CAPLUS [Full-text](#)

DN 78:111557

OREF 78:17915a,17918a

TI Absolute configuration of cryptostylinines I, II, and III by x-ray analysis and aromatic chirality method

AU Blount, J. P.; Toome, V.; Teitel, S.; Brossi, A.

CS Chem. Res. Dep., Hoffman-LaRoche Inc., Nutley, NJ, USA

SO Tetrahedron (1973), 29(1), 31-9

CODEN: TETRAB; ISSN: 0040-4020

DT Journal

LA English

OS CASREACT 78:111557

GI For diagram(s), see printed CA Issue.

AB Single crystal x-ray anal. of unnatural cryptostyline II (I, R = Me) hydrobromide established its absolute configuration as R. Thus, the natural isomer (II, R = H, R1 = R2 = OMe) and the related alkaloids cryptostyline I (II, R = H, R1R2 = OCH2O) and cryptostyline III (II, R = R1 = R2 = OMe) have S configurations. The orthorhombic crystals of I.HBr (R = Me), space group P212121, had a 10.162, b 12.352, c 16.456 Å, d.(observed) 1.37, and d.(calculated) 1.364, for Z = 4. The structure was solved by the heavy atom method and refined by least squares to R 0.031 for 1882 observed reflections. The configuration was confirmed by the aromatic chirality method and by the CD spectrum of the monophenol (I, R = H).

IT 22325-16-2

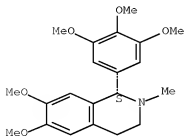
RL: PRP (Properties)

(absolute configuration of)

RN 22325-16-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

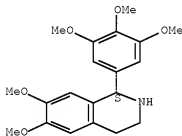
Absolute stereochemistry. Rotation (+).



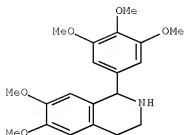
L5 ANSWER 54 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1973:4145 CAPLUS Full-text
 DN 78:4145
 OREF 78:691a,694a
 TI Optically active 1-phenylisoquinolines
 IN Kametani, Tetsuji
 PA Grelan Pharmaceutical Co., Ltd.
 SO Jpn. Kokai Tokkyo Koho, 6 pp.
 CODEN: JKXXAF
 DT Patent
 LA Japanese
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------|----------|-----------------|--------------|
| PI | JP 47025183 | B4 | 19721019 | JP 1971-13124 | 19710312 <-- |
| AB | Optically inactive 1-phenylisoquinolines were treated with optically active salt forming agent so that one of the optically active salts could be selectively crystallized E.g., 1,2,3,4-tetrahydro-o,p-dimethoxy-1-(3,4,5-trimethoxyphenyl)isoquinoline, prepared by reduction of the corresponding 3,4-dihydro compound, was treated with di-p-toluoyl-(+)-tartaric acid in Me2CO and the separated crystals decomposed with Na2CO3 solution to give (-)-compound | | | | |
| IT | 32886-69-4P | | | | |
| | RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of) | | | | |
| RN | 32886-69-4 | CAPLUS | | | |
| CN | Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, (S)- (9CI) (CA INDEX NAME) | | | | |

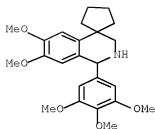
Absolute stereochemistry.



IT 33033-84-0
 RL: PROC (Process)
 (resolution of)
 RN 33033-84-0 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
 (CA INDEX NAME)



L5 ANSWER 55 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1972:72372 CAPLUS Full-text
 DN 76:72372
 OREF 76:11649a,11652a
 TI Isoquinoline derivatives. V. Synthesis of some 1-substituted
 6,7-dimethoxy-1,2,3,4-tetrahydroisoquinoline-4-spirocyclopentanes and
 their analogs
 AU Mndzhoyan, A. L.; Markaryan, E. A.; Arustamyan, Zh. S.; Marashyan, E. S.
 CS Inst. Tonkoi Org. Khim., Erevan, USSR
 SO Khimiya Geterotsiklichesikh Soedinenii (1971), 7(5), 637-40
 CODEN: KGSSAQ; ISSN: 0132-6244
 DT Journal
 LA Russian
 GI For diagram(s), see printed CA Issue.
 AB A series of the title compds. (I, R = H, MeO; R1 = H, MeO; R2 = MeO, H; n = 0,
 1) [prepared by Bischler-Napieralski reaction of the 1-amidomethyl-1-(3,4-
 dimethoxyphenyl)cyclopentane (II, n = 0, 1) in boiling PhMe with POC13 and
 subsequent reduction with LiAlH4 of the 1,2-double bond in the 6,7-dimethoxy-
 3,4-dihydroisoquinoline-4-spirocyclopentane (III, n = 0, 1)], and the 1-
 aminomethyl-1-(3,4- dimethoxyphenyl)cyclopentanes (IV, n = 0, 1) (prepared from
 II by reduction with LiAlH4) in the form of their hydrochlorides produced a
 decrease in blood pressure of about 30 mm of Hg when given intravenously in a
 dose 1-3 mg/kg. II were prepared from the corresponding amines (V) (obtained
 from 3,4-(MeO)2C6H4-CH2CN via alkylation with Br(CH2)4Br in the presence of
 NaNH2 followed by reduction with LiAlH4) and acid chlorides in C6H6 in the
 presence of stoichiometric amts. of pyridine.
 IT 34976-64-2
 RL: PROC (Process)
 (preparation of)
 RN 34976-64-2 CAPLUS
 CN Spiro[cyclopentane-1,4'-(1'H)-isoquinoline],
 2',3'-dihydro-6',7'-dimethoxy-1'-(3,4,5-trimethoxyphenyl)-, hydrochloride
 (1:1) (CA INDEX NAME)



● HCl

OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

L5 ANSWER 56 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1971:529977 CAPLUS [Full-text](#)

DN 75:129977

OREF 75:20531a,20534a

TI Synthesis and absolute configuration of cryptostylinines I, II, and III

AU Brossi, A.; Teitel, S.

CS Chem. Res. Dep., Hoffmann-La Roche Inc., Nutley, NJ, USA

SO Helvetica Chimica Acta (1971), 54(6), 1564-71

CODEN: HCACAV; ISSN: 0018-019X

DT Journal

LA English

GI For diagram(s), see printed CA Issue.

AB The title compds. [I (RR1 = OCH2O, R2 = H; R = R1 = OMe, R2 = H; R = R1 = R2 = OMe)] were prepared by the reductive N-methylation of secondary norcryptostylinines. The natural I possessed the (S) configuration. The reported easy racemization ability of the natural I was traced to optical impurities in the original preparation

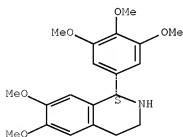
IT 32886-69-4P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation and O.R.D. of)

RN 32886-69-4 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
, (S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



IT 33033-86-2F

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
(preparation and absolute configuration of)

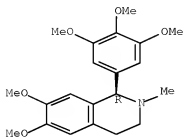
RN 33033-86-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-

10/591,174

trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



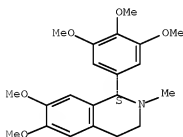
IT 22325-16-2P

RL: PRP (Properties); SPN (Synthetic preparation); PREP (Preparation)
(preparation and absolute configuration of, O.R.D. and)

RN 22325-16-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

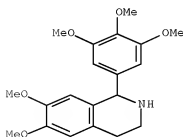


IT 33033-84-0P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation and resolution of)

RN 33033-84-0 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



IT 32886-70-7P 33755-95-2P 33770-51-3P

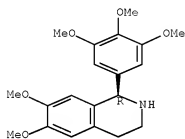
33839-85-6P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 32886-70-7 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
, (R)- (9CI) (CA INDEX NAME)

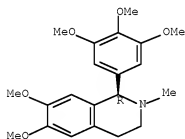
Absolute stereochemistry.



RN 33755-95-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, hydrobromide, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

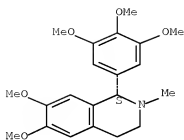


● HBr

RN 33770-51-3 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, hydrobromide (1:1), (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



● HBr

RN 33838-85-6 CAPLUS

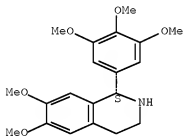
CN L-xylo-Hexulofuranosonic acid, 2,3:4,6-di-O-isopropylidene-, α -,
compd. with (S)-(-)-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-
trimethoxyphenyl)isoquinoline (1:1) (8CI) (CA INDEX NAME)

CM 1

CRN 32886-69-4

CMF C20 H25 N O5

Absolute stereochemistry.



CM 2

CRN 18467-77-1

CMF C12 H18 O7

Absolute stereochemistry. Rotation (-).

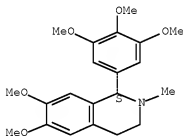


OSC.G 5 THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)

L5 ANSWER 57 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

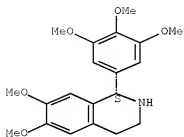
AN 1971:464052 CAPLUS [Full-text](#)
 DN 75:64052
 OREF 75:10159a,10162a
 TI Synthesis of heterocyclic compounds. CCCXCVII. Absolute configuration of cryptostyline. III
 AU Kametani, T.; Sugi, H.; Shibuya, S.
 CS Pharm. Inst., Tohoku Univ., Sendai, Japan
 SO Tetrahedron (1971), 27(12), 2409-14
 CODEN: TETRAB; ISSN: 0040-4020
 DT Journal
 LA English
 GI For diagram(s), see printed CA Issue.
 AB Optical resolution of (\pm)-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)isoquinoline [(\pm)-I] with di-p-toluoyl-(+)-tartaric acid gave (-)-(\pm)-I and (+)-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)isoquinoline. I was transformed to (+)-cryptostyline III (II). A method for the stereochem. study of cryptostyline III by correlation of ORD and CD spectra with those of (+) (R)-1,2,3,4-tetrahydro-6-methoxy-2-methyl-1-phenylisoquinoline.
 IT 22325-16-2
 RL: PRP (Properties)
 (configuration of, absolute)
 RN 22325-16-2 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).



IT 32886-69-3F 32886-69-4P 32886-70-7P
 33033-85-1F 33033-86-2P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)
 RN 32886-68-3 CAPLUS
 CN Butanedioic acid, 2,3-bis[(4-methylbenzoyl)oxy]-, [S-(R*,R*)]-, compd. with (S)-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)isoquinoline (1:2) (9CI) (CA INDEX NAME)
 CM 1
 CRN 32886-69-4
 CMF C20 H25 N O5

Absolute stereochemistry.

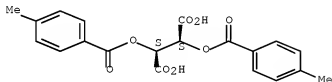


CM 2

CRN 32634-68-7

CMF C20 H18 O8

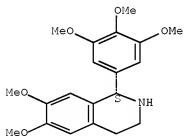
Absolute stereochemistry. Rotation (+).



RN 32886-69-4 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, (S)- (9CI) (CA INDEX NAME)

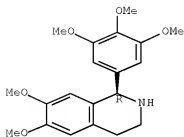
Absolute stereochemistry.



RN 32886-70-7 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

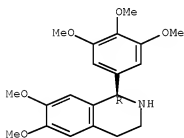


RN 33033-85-1 CAPLUS
 CN Butanedioic acid, 2,3-bis[(4-methylbenzoyl)oxy]-, [R-(R*,R*)]-, compd.
 with (R)-1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-
 trimethoxyphenyl)isoquinoline (1:2) (9CI) (CA INDEX NAME)

CM 1

CRN 32886-70-7
 CMF C20 H25 N O5

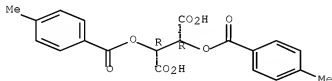
Absolute stereochemistry.



CM 2

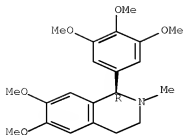
CRN 32634-66-5
 CMF C20 H18 O8

Absolute stereochemistry. Rotation (-).

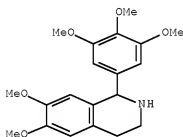


RN 33033-86-2 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-
 trimethoxyphenyl)-, (R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



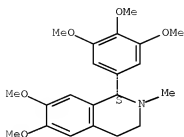
IT 33033-84-0
 RL: PROC (Process)
 (resolution of)
 RN 33033-84-0 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
 (CA INDEX NAME)



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

L5 ANSWER 58 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 1969:106718 CAPLUS [Full-text](#)
 DN 70:106718
 OREF 70:19943a,19946a
 TI Orchidaceae alkaloids. XI. Three
 1-phenyl-1,2,3,4-tetrahydroisoquinolines from *Cryptostylis fulva*
 AU Leander, Kurt; Luning, Bjorn; Ruusa, Ene
 CS Univ. Stockholm, Stockholm, Swed.
 SO Acta Chemica Scandinavica (1947-1973) (1969), 23(1), 244-8
 CODEN: ACSAA4; ISSN: 0001-5393
 DT Journal
 LA English
 AB Three 1-phenyl-2-methyl-6,7-dimethoxy-1,2,3,4-tetrahydroiso-quinolines were
 isolated from *C. fulva*. The 3 alkaloids differ in the substitution pattern of
 the 1-phenyl group, being 3,4-methylenedioxy, 3,4-dimethoxy, and 3,4,5-
 trimethoxy, resp. Their structures were assigned by spectral methods and by
 synthesis.
 IT 22325-16-2
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (new alkaloid from *Cryptostylis fulva*, structure of)
 RN 22325-16-2 CAPLUS
 CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-
 trimethoxyphenyl)-, (1S)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

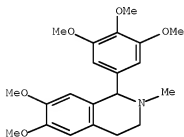


IT 22324-83-0P

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 22324-83-0 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)- (CA INDEX NAME)



OSC.G 5 THERE ARE 5 CAPLUS RECORDS THAT CITE THIS RECORD (5 CITINGS)

L5 ANSWER 59 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1965:93561 CAPLUS [Full-text](#)

DN 62:93561

OREF 62:16783g-h

TI Pharmacological activity of some papaverine analogs. II

AU Sheikova, Zh.; Nikolova, M

SO Izvestiya na Instituta po Fiziologiya, Bulgarska Akademiya na Naukite (1964), 7, 243-52

CODEN: IIFBA4; ISSN: 0068-3922

DT Journal

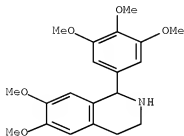
LA Bulgarian

AB cf. CA 60, 14672h. The spasmolytic activity of the nicotinoyl, isonicotinoyl, and furanoyl derivs. were lower than papaverine (I). The o-methoxyphenol and trimethoxyphenol derivs. had equal or superior spasmolytic effects. The nonhydrated compds. of these derivs. had a depressive effect on the central nervous system, while the hydrated compds. had a stimulant effect. All compds., except those containing furanoyl and o-methoxyphenyl groups were less toxic than I.

IT 33033-84-0, Isoquinoline,
1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
(antispasmodic activity of)

RN 33033-84-0 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
(CA INDEX NAME)



L5 ANSWER 60 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1965:93560 CAPLUS [Full-text](#)

DN 62:93560

OREF 62:16783e-g

TI Relations between chemical structure and pharmacological actions in a series of γ -lactones and amino alcohols

AU Zavrazhnov, V. I.; Ponomarev, F. G.; Trukhacheva, L. I.

CS Med. Inst., Voronezh

SO Elektron. i Khim. v Kardiolog., Voronezhsk. Obl. Obshchestvo Kardiologov, Voronezhsk. Obl. Obshchestvo Terapevtov (1964) 348-58

DT Journal

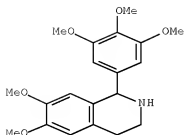
LA Russian

AB The effects were investigated of synthetic derivs. of α -aceto- γ -butyrolactone (I), Et acetoacetate, morpholine (II) derivs., and derivs. of diethylaminopropanediol ether (III) on perfused blood vessels of frog's hind limb and of isolated rabbit ear, on rabbit intestinal strips in vitro, and on rabbit blood pressure. The L.D.100 was determined in mice. The introduction of Me and vinyl groups in the β -position or in both β - and γ -positions of I caused a reversal of its pharmacol. properties: the derivs. have a spasmolytic activity on blood vessels without any influence on systemic blood pressure and intestinal smooth muscle. Methylation in the β -position or in both β - and γ -positions enhanced the vasopressor activity of I. The introduction of MeOCH₂, PrOCH₂, and iso-PrOCH₂ groups into I in the γ -position caused an increase of pressor properties; substitution of the propoxy group in the γ -position with Cl decreased the pressor activity of the compound and toxicity but had a neg. influence on intestinal peristaltic movements. Substitution of H. in the NH group of II derivs. by aliphatic groups caused an increase of pressor properties and toxicity. From 7 derivs. of III the most active ones were benzyl and Et derivs. which decreased rabbit blood pressure and caused a constriction of ear and limb blood vessels. All III derivs. inhibited the movements and tonus of the rabbit intestine.

IT 33033-84-0, Isoquinoline,
1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
(antispasmodic activity of)

RN 33033-84-0 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-1-(3,4,5-trimethoxyphenyl)-
(CA INDEX NAME)



L5 ANSWER 61 OF 61 CAPLUS COPYRIGHT 2009 ACS on STN

AN 1955:84276 CAPLUS [Full-text](#)

DN 49:84276

OREF 49:15902h-i,15903a-i

TI Chemistry of vanillin and its derivatives. VI. Effective spasmolytic 1-phenylisoquinolines and diphenethylamines containing the guaiacyl grouping

AU Kratzl, K.; Horejschi, T.; Billek, G.

CS Univ. Vienna

SO Monatshefte fuer Chemie (1954), 85, 1154-65

CODEN: MOCMB7; ISSN: 0026-9247

DT Journal

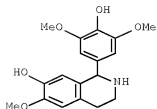
LA Unavailable

OS CASREACT 49:84276

AB cf. C.A. 48, 1363g. Several 1-phenylisoquinolines were prepared from 4-(2-aminoethyl)guaiacol (I) and its benzyl ether (II), by forming amides, treating these with POCl₃, and dehydrogenating. The following amides were prepared by treatment with the corresponding acyl or alkyl chloride (starting material, yield, and m.p. given): BzNHCH₂CH₂C₆H₃(OH)OMe-4,3, I, 94, 127°; BzNHCH₂CH₂C₆H₃(OBz)OMe-4,3 (III), I, -, -; BzNHCH₂CH₂C₆H₃(OCH₂Ph)OMe-4,3 (IV), I (or II), 76, 134°; 3,4-MeO(AcO)C₆H₃CONHCH₂CH₂C₆H₃(OH)OMe-4,3, I, 89, 117-20°; 3,4-MeO(BzO)C₆H₃CONHCH₂CH₂C₆H₃(OH)OMe-4,3 (V), I, 95, 133-6°; 3,4-MeO(PhCH₂O)C₆H₃CONHCH₂CH₂C₆H₃(OH)OMe-4,3 (VI), I, 93, 146-7°; 3,4-MeO(BzO)C₆H₃CONHCH₂CH₂C₆H₃(OBz)OMe-4,3, V, 87, 154-5°; 3,4-MeO(PhCH₂O)C₆H₃CONHCH₂CH₂C₆H₃(OBz)OMe-4,3, VI, 90, 143°; 3,4-MeO(PhCH₂O)C₆H₃CONHCH₂CH₂C₆H₃(OCH₂Ph)OMe-4,3 (VII), IV or II, 75-89, 167°; and 3,5,4-(MeO) 2 (PhCH₂O)C₆H₃CONHCH₂CH₂C₆H₃(OCH₂Ph)OMe-4,3 (VIII), II, 67, 125°. For the ring-closure, 5 millimoles of the amide was treated with 12.5 millimoles POCl₃ in 60 ml. absolute xylene or PhMe at reflux for 20-60 min. The product was precipitated as a resinous mass by addition of petr. ether. The free base could be precipitated from dilute HCl solution (containing EtOH if necessary) by NH₃ and recrystd. from EtOH-H₂O. The hydrochloride could be recovered instead by excess concentrated HCl. The picrate was prepared from the free base with picric acid or from the hydrochloride with sodium picrate. In this way were made the following 3,4-dihydroisoquinolines and derivs. (substituents, starting compound, % yield, m.p. given): 7-benzoyloxy-6-methoxy-1-phenyl (IX), III, 78, 191-2° (picrate, m. 210-12°; hydrochloride, m. 209-12°); 7-benzoyloxy-6-methoxy-1-phenyl, IV, 88, 137° (hydrochloride, m. 199°); 7-benzoyloxy-1-(4-benzoyloxy-3-methoxyphenyl)-6-methoxy (X), VII, 79, - (picrate, m. 203°); 7-benzoyloxy-1-(4-benzoyloxy-3,5-dimethoxyphenyl)-6-methoxy (XI), VIII, 77, - (picrate, m. 232°). Hydrolysis of IX with aqueous EtOH-NaOH gave 93% 6-methoxy-1-phenyl-3,4-dihydro-7-isoquinolinol, m. 180° (picrate, m. 240°; hydrochloride, m. 210-12°). Treatment of X with 20% HCl and recrystn. from dilute HCl gave 93% 1-(4-hydroxy-3-methoxyphenyl)-6-methoxy-3,4-dihydro-7-isoquinolinol hydrochloride (m. 258°). Similarly, XI gave 92% 1-(4-hydroxy-3,5-dimethoxyphenyl)-6-methoxy-3,4-dihydro-7-isoquinolinol hydrochloride (m.

234°, with 1 mol. H₂O). The free bases (XII and XIII) could be obtained with K₂CO₃. Hydrogenation and recrystallization from EtOH or dilute HCl gave from XII 94% 1-(4-hydroxy-3-methoxyphenyl)-6-methoxy-1,2,3,4-tetrahydro-7-isoquinolinol hydrochloride (m. 182°), and from XIII 92% 1-(4-hydroxy-3,5-dimethoxyphenyl)-6-methoxy-1,2,3,4-tetrahydro-7-isoquinolinol hydrochloride (m. 150°, with 1 mol. H₂O). The free bases (XIV and XV) were obtained by K₂CO₃ treatment. Pd dehydrogenation gave from XI 90% 7-benzoyloxy-6-methoxy-1-phenylisoquinoline (XVI) (oil); from XIV 50% 1-(4-hydroxy-3-methoxyphenyl)-6-methoxy-7-isoquinolinol (hydrochloride, from EtOH-2N HCl, m. 240°; picrate, m. 220°); and from XV 56% 1-(4-hydroxy-3,5-dimethoxyphenyl)-6-methoxy-7-isoquinolinol (hydrochloride, from dilute HCl, m. 217°; picrate, m. 242-3°). Hydrolysis of XVI with aqueous EtOH-NaOH gave 7% 6-methoxy-1-phenyl-7-isoquinolinol (hydrochloride, m. 145-7°; picrate, m. 221-2°). 4-Benzoyloxy-3,5-dimethoxybenzoic acid (XVII) (55.6%, m. 155-7° from EtOH-H₂O) was prepared from 2 g. syringic acid with PhCH₂Cl and KOH in EtOH. XVII formed 4-benzoyloxy-3,5-dimethoxybenzoyl chloride (85%, m. 45°) with SOCl₂ at 50-70°. Syringaldehyde (XVIII) (48%, m. 110-12°, from H₂O) was prepared from 6.2 g. 5-methoxyprotocatechualdehyde with 12.6 g. Me₂SO₄ in 40 ml. H₂O containing 9.3 g. NaOH, then acidification with HCl. XVIII sodium salt gave 62.5% 4,3,5-PhCH₂O(MeO)2C₆H₃CHO (XIX) (m. 63° from EtOH) with PhCH₂Cl in xylene. XIX (3.2 g.) in 25 ml. absolute EtOH with 0.8 g. MeNO₂, 0.13 g. MeNH₂.HCl, and 0.1 g. Na₂CO₃ was heated 24 hrs. at 40° to give 65% 4,3,5-PhCH₂O(MeO)₂, C₆H₃CH:CHNO₂ (m. 133° from C₆H₆-EtOH). [3,4-MeO(PhCH₂O)C₆H₃]2NH₂.HCl (XX) (m. 205-10° from EtOH) was prepared two ways: (A) 0.5 g. 3,4-MeO(PhCH₂O)C₆H₃CH:CHNO₂ was dissolved in 17 ml. HOAc plus 12 ml. C₆H₆. The solution was dropped into 50 mg. PtO₂ in 3 ml. HOAc under H. After taking up 150 ml. H, the solution was filtered and evaporated to 20 ml. under reduced pressure. Addition of concentrated HCl precipitated 51% XX; (B) 1 g. 3,4-MeO(PhCH₂O)C₆H₃CH₂CH:NOH was dissolved in 35 ml. EtOH and added to 50 mg. PtO₂ in 5 ml. EtOH under H. After absorption of 165 ml. H, XX (38%) was recovered as in A. XX (1 g.) was heated 3 hrs. at 130-40° with 14 ml. 38% formalin, the mixture was cooled and 2N HCl added to give 80% 4,4'-bisbenzyloxy-3,3'-dimethoxy-N-methylidiphenethylamine hydrochloride (m. 176-8°, from EtOH containing concentrated HCl; 1/2 mol. H₂O of crystallization). XX (0.5 g.) on hydrolysis with 10 ml. concentrated HCl in 10 ml. EtOH gave 88% 4,4'-iminodihydroxydiguaiacol hydrochloride (m. 205-8°, from concentrated HCl). All the isoquinolines, phenethylamines, and intermediates showed 1/5 to 1/20 the spasmolytic activity of papaverine.

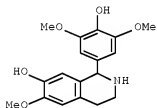
IT 855737-16-5P, 7-Isoquinolinol,
1,2,3,4-tetrahydro-1-(4-hydroxy-3,5-dimethoxyphenyl)-6-methoxy-,
hydrochloride 855737-16-7P, 7-Isoquinolinol,
1,2,3,4-tetrahydro-1-(4-hydroxy-3,5-dimethoxyphenyl)-6-methoxy-
RL: PREP (Preparation)
(preparation of)
RN 855737-16-5 CAPLUS
CN 7-Isoquinolinol, 1,2,3,4-tetrahydro-1-(4-hydroxy-3,5-dimethoxyphenyl)-6-methoxy-, hydrochloride (1:1) (CA INDEX NAME)



● HCl

RN 855737-18-7 CAPLUS

CN 7-Isoquinolinol, 1,2,3,4-tetrahydro-1-(4-hydroxy-3,5-dimethoxyphenyl)-6-methoxy- (CA INDEX NAME)



=> s 14 not 15

L6 14 L4 NOT L5

=> dis l6 1-14 bib abs fhitrstr

L6 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2007:703811 CAPLUS Full-text

DN 147:118219

TI Preparation of isoquinoline aminopyrazole derivatives for the treatment of cancer

IN Chen, Li; Georges, Guy; Mertens, Alfred; Wu, Xihan

PA F. Hoffmann-La Roche AG, Switz.

SO PCT Int. Appl., 334pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|----------|-----------------|----------|
| PI WO 2007071348 | A1 | 20070628 | WO 2006-EP12112 | 20061215 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, | | | | |

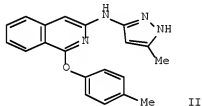
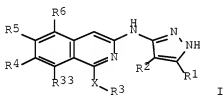
CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
KG, KZ, MD, RU, TJ, TM

| | | | | |
|----------------------------------------------------------------------------------------------------------------------------------|----|----------|------------------|----------|
| US 20070179151 | A1 | 20070802 | US 2006-637733 | 20061212 |
| US 7572809 | B2 | 20090811 | | |
| AU 2006328997 | A1 | 20070628 | AU 2006-328997 | 20061215 |
| CA 2633101 | A1 | 20070628 | CA 2006-2633101 | 20061215 |
| EP 1966190 | A1 | 20080910 | EP 2006-840991 | 20061215 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR | | | | |
| JP 2009519918 | T | 20090521 | JP 2008-544887 | 20061215 |
| NO 2008002643 | A | 20080901 | NO 2008-2643 | 20080605 |
| MX 2008007624 | A | 20080701 | MX 2008-7624 | 20080612 |
| ZA 2008005163 | A | 20090624 | ZA 2008-5163 | 20080612 |
| KR 2008076963 | A | 20080820 | KR 2008-714869 | 20080619 |
| CN 101341145 | A | 20090107 | CN 2006-80048040 | 20080619 |
| IN 2008CN03084 | A | 20090306 | IN 2008-CN3084 | 20080619 |
| PRAI EP 2005-27720 | A | 20051219 | | |
| WO 2006-EP12112 | W | 20061215 | | |

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

OS MARPAT 147:118219

GI



AB The title compds. I [R1 = H, alkyl, cycloalkyl; R2 = H or alkyl; R3 = (un)substituted alkyl, aryl, heteroaryl, etc.; R33 = H, alkyl or alkoxy; R4 = H, alkyl, alkoxy, etc.; R5 = H, alkyl, alkoxy, etc.; R6 = H, alkyl, alkoxy, etc. or R5 and R6 can form together with the carbon atoms to which they are attached a 5-6 membered heterocyclic ring; X = a bond, O, S, C(O), etc.], useful in the control or prevention of illnesses such as cancer, were prepared and formulated. E.g., a multi-step synthesis of II, starting from 1-indanone, was given. II showed IC50 of 0.066 μ M against Aurora A kinase.

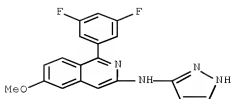
IT 942932-52-7P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of isoquinolinylamino pyrazoles for the treatment of cancer)

RN 942932-52-7 CAPLUS

CN 3-Isoquinolinamine, 1-(3,5-difluorophenyl)-6-methoxy-N-1H-pyrazol-3-yl- (CA INDEX NAME)



OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
 RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2007:565283 CAPLUS Full-text
 DN 146:528334
 TI Novel colored solutions of injectable drugs and their salts
 IN Winch, Peter D.
 PA Winch, Peter, D., USA
 SO PCT Int. Appl., 121 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|-----------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------|------------------|----------|
| PI | WO 2007059019 | A2 | 20070524 | WO 2006-US43963 | 20061113 |
| | WO 2007059019 | A3 | 20080710 | | |
| | W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | |
| | RW: | AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AP, EA, EP, OA | | | |
| | AU 2006315610 | A1 | 20070524 | AU 2006-315610 | 20061113 |
| | CA 2635496 | A1 | 20070524 | CA 2006-2635496 | 20061113 |
| | EP 1951248 | A2 | 20080806 | EP 2006-837427 | 20061113 |
| | R: | AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, RS | | | |
| | JP 2009515895 | T | 20090416 | JP 2008-540243 | 20061113 |
| | CN 101400369 | A | 20090401 | CN 2006-80050729 | 20080709 |
| | US 20090156562 | A1 | 20090618 | US 2008-93534 | 20081118 |
| PRAI | US 2005-735370P | P | 20051114 | | |
| | US 2005-736372P | P | 20051114 | | |
| | US 2005-736373P | P | 20051114 | | |
| | US 2005-736374P | P | 20051114 | | |
| | US 2005-736464P | P | 20051114 | | |
| | US 2005-736468P | P | 20051114 | | |
| | US 2005-736573P | P | 20051114 | | |

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|-----------------|---|----------|
| US 2005-736574P | P | 20051114 |
| US 2005-736575P | P | 20051114 |
| US 2005-736576P | P | 20051114 |
| US 2005-736577P | P | 20051114 |
| US 2005-736578P | P | 20051114 |
| US 2005-736579P | P | 20051114 |
| US 2006-761274P | P | 20060123 |
| US 2006-761276P | P | 20060123 |
| US 2006-761277P | P | 20060123 |
| US 2006-761282P | P | 20060123 |
| US 2006-761283P | P | 20060123 |
| WO 2006-US43963 | W | 20061113 |

AB The invention is directed to pharmaceutical compns. comprising colored solns., colored emulsions, or colored powders of injectable pharmaceuticals wherein the pharmaceuticals are selected from the group consisting of muscle relaxants, hypnotics, induction agents, and anticholinergics. The formulations of the present invention may all be colored using fluorescein. Different colors may be achieved by either varying the concentration of fluorescein, or by combining fluorescein with another dye. The invention is also directed to methods involving the use of said pharmaceutical compns. A sterile aqueous solution contains atracurium besylate 10 mg/mL, and fluorescein (10 mg), wherein the pH of the solution is adjusted to 3.5 with benzenesulfonic acid.

IT 213998-46-0

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(colored solns. of injectable drugs and their salts)

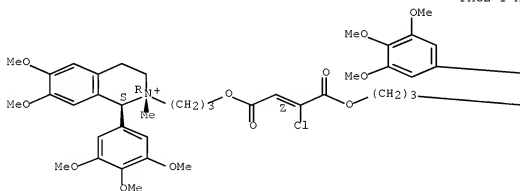
RN 213998-46-0 CAPLUS

CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

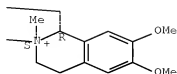
Absolute stereochemistry.

Double bond geometry as shown.

PAGE 1-A



● 2 Cl⁻



L6 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2009 ACS ON STN

AN 2007:284437 CAPLUS Full-text

DN 146:337745

TI Preparation of tetrahydroisoquinoline derivatives as IGF-1 receptor inhibitors

IN Gunzinger, Jan; Leander, Kurt

PA Analytecon SA, Switz.

SO PCT Int. Appl., 52pp.

CODEN: PIXXD2

DT Patent

LA English

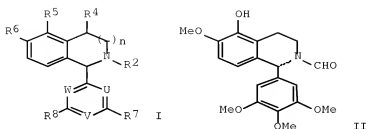
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|----------|------------------|----------|
| PI | WO 2007029107 | A1 | 20070315 | WO 2006-IB2474 | 20060908 |
| | W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| | RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| | AU 2006288847 | A1 | 20070315 | AU 2006-288847 | 20060908 |
| | CA 2621820 | A1 | 20070315 | CA 2006-2621820 | 20060908 |
| | EP 1940796 | A1 | 20080709 | EP 2006-795449 | 20060908 |
| | R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, HR, RS | | | | |
| | JP 2009507820 | T | 20090226 | JP 2008-529711 | 20060908 |
| | ZA 2008001910 | A | 20090826 | ZA 2008-1910 | 20080229 |
| | MX 2008003298 | A | 20080411 | MX 2008-3298 | 20080307 |
| | NO 2008001206 | A | 20080529 | NO 2008-1206 | 20080307 |
| | CN 101282935 | A | 20081008 | CN 2006-80032919 | 20080307 |
| | KR 2008065591 | A | 20080714 | KR 2008-706725 | 20080320 |
| | IN 2008DN02917 | A | 20080704 | IN 2008-DN2917 | 20080408 |
| | US 20090099133 | A1 | 20090416 | US 2008-991531 | 20080415 |
| PRAI | US 2005-715170P | P | 20050909 | | |
| | WO 2005-IB2667 | A | 20050909 | | |
| | WO 2006-IB2474 | W | 20060908 | | |

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

OS CASREACT 146:337745; MARPAT 146:337745

GI



AB The title compds. with general formula I [wherein R₄ = H, OH, CN, CF₃, etc.; R₂ = H, Me, Et, etc.; R₅ = H, alkyl, OH, etc.; R₆ = Me, halo, alkoxy, etc.; n = 1-2; R₇ and R₈ = independently OH, Me, Et, methoxy, etc.; U = N or CR₁, where R₁ = H, alkyl, alkoxy, etc.; V = N or CR₃, where R₃ = H, OH, halo, etc.; W = N or CR₉, where R₉ = H, halo, CF₃, etc.] or pharmaceutically acceptable salts thereof are prepared as inhibitors of IGF-1 receptor. For example, compound II and its prodrug, (1R)-1-(3,4,5-trimethoxyphenyl)-2-formyl-5-(dihydrogen phosphate)-6-methoxy-1,2,3,4-tetrahydroisoquinoline, were prepared from a multi-step synthesis. II showed IC₅₀ values of 50 nM and 40 nM for the inhibition of phospho-MAPK (Erk1/2) and phospho-AKT, resp. I, as inhibitors of IGF-1R, are useful in the treatment of diseases such as but not limited to cancer, atherosclerosis, and psoriasis (no data).

IT 929050-66-8P

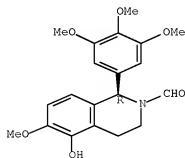
RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(drug candidate; preparation of tetrahydroisoquinoline derivs. as IGF-1 receptor inhibitors)

RN 929050-66-8 CAPLUS

CN 2(1H)-Isoquinolinecarboxaldehyde, 3,4-dihydro-5-hydroxy-6-methoxy-1-(3,4,5-trimethoxyphenyl)-, (1R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

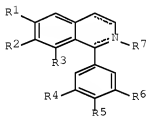
AN 2007:64394 CAPLUS [Full-text](#)

DN 146:229193

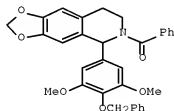
TI Preparation of isoquinoline derivatives for treatment of tumor

IN Zhao, Yu; Ding, Hongxia; Lu, Wei
 PA Zhejiang University, Peop. Rep. China
 SO Faming Zhuanli Shenqing Gongkai Shuomingshu, 38 pp.
 CODEN: CNXXEV
 DT Patent
 LA Chinese
 FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|----------------------------------------|------|----------|------------------|----------|
| PI | CN 1896065 | A | 20070117 | CN 2005-10082721 | 20050711 |
| | CN 101440096 | A | 20090527 | CN 2008-10189046 | 20050711 |
| | CN 101440097 | A | 20090527 | CN 2008-10189066 | 20050711 |
| PRAI | CN 2005-10082721 | A3 | 20050711 | | |
| OS | CASREACT 146:229193; MARPAT 146:229193 | | | | |
| GI | | | | | |



I



II

AB The title 1-(3',4',5'-trisubstituted phenyl)isoquinoline derivs. I [R1-R3 = independently H, alkoxy, benzyloxy, hydroxy, halo, or amino; or R1 and R2 = -OCH2O- or -OCH2CH2O-; R4-R6 = independently alkoxy, benzyloxy, hydroxy, halo, or amino; or R4 and R5 or R5 and R6 = -OCH2O- or -OCH2CH2O-; R7 = absence, H, (un)substituted benzoyl, or cyclopropylcarbonyl; with provisos], or pharmaceutically acceptable salts, solvates, or mixts. thereof were prepared for the treatment of tumor (no data). For example, II was prepared in a multi-step synthesis. The compds. showed excellent antitumor activity.

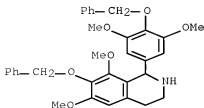
IT 903527-19-5P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

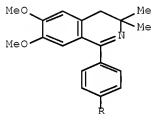
(drug candidate; preparation of isoquinoline derivs. for treatment of tumor)

RN 903527-19-5 CAPLUS

CN Isoquinoline, 1-[3,5-dimethoxy-4-(phenylmethoxy)phenyl]-1,2,3,4-tetrahydro-6,8-dimethoxy-7-(phenylmethoxy)- (CA INDEX NAME)



L6 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2006:1238764 CAPLUS [Full-text](#)
 DN 146:229148
 TI Synthesis and anticoagulant activity of 1-aryl derivatives of tetrahydroisoquinolines
 AU Glushkov, V. A.; Arapov, K. A.; Minova, O. N.; Ismailova, N. G.; Syropyatov, B. Ya.; Shklyayev, Yu. V.
 CS Perm State University, Perm, Russia
 SO Pharmaceutical Chemistry Journal (2006), 40(7), 363-366
 CODEN: PCJOAU; ISSN: 0091-150X
 PB Springer
 DT Journal
 LA English
 OS CASREACT 146:229148
 GI



I

AB A series of 1-aryl-3,3-dimethyl-3,4-dihydroisoquinolines, e.g., I (R = H, Br, NO₂ or MeO), were obtained by three-component (one-pot) condensation of veratrole, isobutylene oxide, and aromatic nitriles and then reduced to the corresponding 1,2,3,4-tetrahydroisoquinolines. Hydrochlorides of the synthesized compds. were tested for anticoagulant activity.

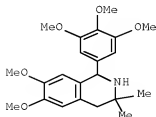
IT 924910-76-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation and anticoagulant activity of aryl dihydro- and tetrahydroisoquinoline derivs. via three-component heterocyclization of veratrole with isobutylene oxide and aromatic nitriles followed by reduction with hydrides)

RN 924910-76-9 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6,7-dimethoxy-3,3-dimethyl-1-(3,4,5-trimethoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME)



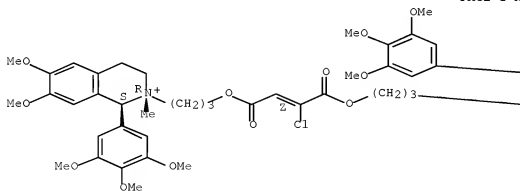
● HCl

OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)
 RE.CNT 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

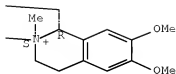
L6 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2006:679649 CAPLUS Full-text
 DN 146:287203
 TI GW280430A (AV430A), a new ultrashort-acting nondepolarizing neuromuscular blockers
 AU Xu, Ji-hong; Zheng, Si-ju
 CS Department of Anesthesiology, General Hospital of Shenyang Command, Shenyang, 110016, Peop. Rep. China
 SO Guowai Yixue Mazuixue Yu Fusu Fence (2005), 26(5), 295-297
 CODEN: GYMYAS; ISSN: 1001-1005
 PB Guowai Yixue Mazuixue Yu Fusu Fence Bianjibu
 DT Journal; General Review
 LA Chinese
 AB A review. GW280430A had such characteristics as rapid effects, short action time, no accumulation and few bad responses, so it was the most promising non-depolarization muscle relaxing medicine which could replace succinylcholine. In this paper GW280430A (AV430A), a new ultrashort-acting nondepolarizing neuromuscular blockers is discussed.
 IT 213998-46-0, GW280430A
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (GW280430A (AV430A), new ultrashort-acting nondepolarizing neuromuscular blockers)
 RN 213998-46-0 CAPLUS
 CN Isoquinolinium, 2-[3-[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.

PAGE 1-A

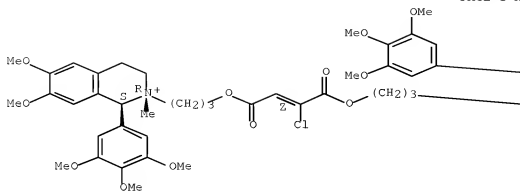
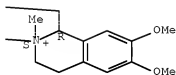


● 2 Cl-



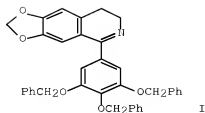
L6 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2005:1309229 CAPLUS Full-text
 DN 144:304254
 TI Muscle relaxants 2006: A clinical and basic science update and commentary
 AU Lee, Chingmuh; Katz, Ronald L.
 CS Department of Anesthesiology, Harbor-UCLA Medical Center, Torrance, CA, USA
 SO Seminars in Anesthesia, Perioperative Medicine and Pain (2005), 24(3), 154-164
 CODEN: SAPPFJ; ISSN: 1547-9951
 PB Elsevier Inc.
 DT Journal; General Review
 LA English
 AB A review. As muscle relaxants remain a mainstay of modern anesthesia practice, it behooves the anesthesiologists to keep themselves up-to-date on the theory and clin. practice of neuromuscular pharmacol. Progress continues to be made in the basic and clin. aspects of neuromuscular pharmacol., including mechanism of action, blocking drugs, and reversal agent. The new mechanism of action is based on the mol. shape of the relaxants. Although the so-called "ideal relaxant" is still not in sight, and may never be, the new relaxant AV430A and the new reversal agent Org 25969 hold potential to significantly improve patient care. AV430 has superior clin. profile, although it is still considerably slower and longer in action than succinylcholine. The fast onset of rocuronium combined with its complete and immediate reversibility with Org 25969 may match succinylcholine in onset and offset. Both drugs are undergoing clin. trials.
 IT 213998-46-0, GW280430A
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (AV430A; AV430 has superior clin. profile but is slower in action than succinylcholine and fast onset of rocuronium combined with complete reversibility with reversal agent Org 25969 may match succinylcholine in onset and offset in patient)
 RN 213998-46-0 CAPLUS
 CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.

● 2 Cl⁻

RE.CNT 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN
AN 2005:1082676 CAPLUS [Full-text](#)
DN 145:188599
TI Synthesis and biological evaluation of novel compounds related to
1-arylnaphthalene lignans and isoquinolines
AU Ding, Hongxia; Lu, Wei; Li, Haibo; Yang, Leixiang; Zhang, Qijun; Zhou,
Changxin; Wu, Xiumei; Baudoin, Olivier; Cai, Junchao; Gueritte, Francoise;
Zhao, Yu
CS Department of Traditional Chinese Medicine and Natural Drug Research,
College of Pharmaceutical Sciences, Zhejiang University, Hangzhou, 310031,
Peop. Rep. China
SO Chemistry & Biodiversity (2005), 2(9), 1217-1231
CODEN: CBHIAM; ISSN: 1612-1872
PB Verlag Helvetica Chimica Acta AG
DT Journal
LA English
OS CASREACT 145:188599
GI



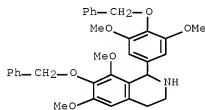
AB Novel compds., e.g. I, designed as hybrids of 1-arylnaphthalene lignans with isoquinoline alkaloids were prepared and evaluated for their cytotoxicities on human tumor cell lines, such as A549, HeLa, PC-3, CNE, BEL-7404, and KB. Some of the synthetic compds. exhibited their IC₅₀ values on selected cell lines at 10⁻⁶ M scale. The preliminary CoMFA mol. modeling studies of these synthetic analogs were also performed.

IT 903527-19-5P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)
(preparation and antitumor activity of aryl-naphthalene lignan-isoquinoline hybrids)

RN 903527-19-5 CAPLUS

CN Isoquinoline, 1-[3,5-dimethoxy-4-(phenylmethoxy)phenyl]-1,2,3,4-tetrahydro-6,8-dimethoxy-7-(phenylmethoxy)- (CA INDEX NAME)



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)
RE.CNT 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2005:1026936 CAPLUS [Full-text](#)

DN 143:326227

TI Preparation of tetrahydroisoquinoline and tetrahydrobenzazepine derivatives as IGF-1R inhibitors

IN Gunzinger, Jan; Leander, Kurt

PA Analytecon S. A., Switz.

SO PCT Int. Appl., 55 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|----|--------------------------------------------------------------------|------|----------|-----------------|----------|
| PI | WO 2005087743 | A1 | 20050922 | WO 2004-CH147 | 20040312 |
| | W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, | | | | |

CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

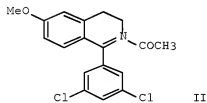
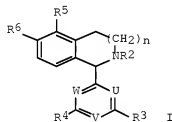
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|---------------|----|----------|-----------------|----------|
| AU 2004317166 | A1 | 20050922 | AU 2004-317166 | 20040312 |
| CA 2555745 | A1 | 20050922 | CA 2004-2555745 | 20040312 |
| EP 1732898 | A1 | 20061220 | EP 2004-719900 | 20040312 |
| EP 1732898 | B1 | 20080123 | | |

R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LI, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, LT, LV, MK

| | | | | |
|--------------------|----|----------|------------------|----------|
| CN 1922147 | A | 20070228 | CN 2004-80042030 | 20040312 |
| BR 2004018462 | A | 20070605 | BR 2004-18462 | 20040312 |
| JP 2007528877 | T | 20071018 | JP 2007-502162 | 20040312 |
| AT 384700 | T | 20080215 | AT 2004-719900 | 20040312 |
| ES 2299825 | T3 | 20080601 | ES 2004-719900 | 20040312 |
| NO 2006003794 | A | 20061212 | NO 2006-3794 | 20060824 |
| US 20070129399 | A1 | 20070607 | US 2006-591174 | 20060830 |
| MX 2006010410 | A | 20061110 | MX 2006-10410 | 20060912 |
| IN 2006DN05643 | A | 20070831 | IN 2006-DN5643 | 20060927 |
| PRAI WO 2004-CH147 | A | 20040312 | | |

OS CASREACT 143:326227; MARPAT 143:326227

GI



AB Title compds. I [R2 = H, Me, Et, etc.; R5 = H, alkyl, OH, etc.; R6 = Me, alkoxy, OCF3, etc.; n = 1-2; R3 and R4 independently = OH, OMe, halo, etc.; U = N or CR1; R1 = H, alkyl, alkoxy, etc.; V = N or CR7; R7 = H, OH, halo, etc.; W = N or CR8; R8 = H, alkyl, alkoxy, etc.] and their pharmaceutically acceptable salts, are prepared and disclosed as inhibitors of IGF-1R. Thus, e.g., II was prepared by amidation of 3-methoxyphenylethylamine with 3,5-dichlorobenzoyl chloride followed by cyclization with POC13 and subsequent reduction to the resp. secondary amine which was then acetylated. The

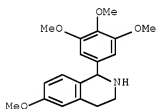
activity of I was evaluated in a cell growth inhibition study using human cell lines MCF-7 and SK-MEL 28 and it was revealed that compds. of the invention possessed IC50 values in the range of 8 µg/mL up to 150 pg/mL in at least one cell line. I as inhibitors of IGF-1R should prove useful in the treatment of diseases such as but not limited to cancer, atherosclerosis and psoriasis. Pharmaceutical compns. comprising I are disclosed.

IT 865151-32-2P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
(preparation of tetrahydroisoquinoline and tetrahydrobenzazepine derivs. as IGF-1R inhibitors)

RN 865151-32-2 CAPLUS

CN Isoquinoline, 1,2,3,4-tetrahydro-6-methoxy-1-(3,4,5-trimethoxyphenyl)-
(CA INDEX NAME)



OSC.G 2 THERE ARE 2 CAPLUS RECORDS THAT CITE THIS RECORD (2 CITINGS)

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2005:715613 CAPLUS Full-text

DN 143:431761

TI Muscle relaxants

AU Takeda, Junzo

CS School of Medicine, Keio University, Japan

SO Sentan Iryo Shirizu (2005), 33(Masuika no Atarashii Nagare), 70-73
CODEN: SISEBJ

PB Sentan Iryo Gijutsu Kenkyusho

DT Journal; General Review

LA Japanese

AB A review, discussing the action mechanism, pharmacokinetics, and pharmacol. of new muscle relaxants, including rocuronium bromide, GW280430A, and TAAC3.

IT 213998-46-6, GW280430A

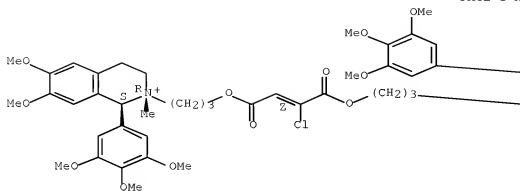
RL: DMA (Drug mechanism of action); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(action mechanism, pharmacokinetics, and pharmacol. of new muscle relaxants, including rocuronium bromide, GW280430A, and TAAC3)

RN 213998-46-0 CAPLUS

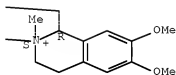
CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.

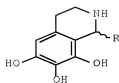
Double bond geometry as shown.



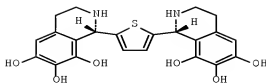
● 2 Cl⁻



L6 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2005:540969 CAPLUS [Full-text](#)
 DN 144:390714
 TI Identification of novel anthrax lethal factor inhibitors generated by
 combinatorial Pictet-Spengler reaction followed by screening in situ
 AU Numa, Mehdi M. D.; Lee, Lac V.; Hsu, Che-Chang; Bower, Kristen E.; Wong,
 Chi-Huey
 CS Department of Chemistry and the Skaggs Institute for Chemical Biology, The
 Scripps Research Institute, La Jolla, CA, 92037, USA
 SO ChemBioChem (2005), 6(6), 1002-1006
 CODEN: CBCHFX; ISSN: 1439-4227
 PB Wiley-VCH Verlag GmbH & Co. KGaA
 DT Journal
 LA English
 OS CASREACT 144:390714
 GI



I



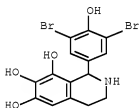
II

AB Anthrax lethal factor (LF) is a zinc-dependent metalloprotease involved in the rapid development of the deadly infection caused by *Bacillus anthracis*. Blocking its action is a plausible method to mitigate the deleterious effects of late stage infection. We report the inhibition of LF by tetrahydroisoquinoline poly phenolic compds. (I) (R = H, COMe, CH(SH)CH₂OH, 2-cyclohexen-1-yl, 2-methylpropyl, n-heptyl, iso-Pr, chloromethyl, etc.), in particular (II), which were identified by screening a combinatorial library of 69 compds. that was generated by Pictet-Spengler reaction of 5-hydroxydopamine hydrochloride with aldehydes or ketones. II inhibited LF with *K_i* of 1.8 μ M under physiol. salt concentration. We also report the identification of com. available polyphenolic inhibitors against LF, e.g., Anthracene Brown G, Resorcin blue, tannic acid, and 1,2,3,4,6-penta-O-galloyl- β -D-glucopyranose.

IT 881426-40-0P
 RL: BSU (Biological study, unclassified); CPN (Combinatorial preparation); BIOL (Biological study); CMBI (Combinatorial study); PREP (Preparation) (identification of anthrax lethal factor inhibitors generated by combinatorial Pictet-Spengler reaction of hydroxydopamine with aldehydes or ketones followed by screening in situ)

RN 881426-40-0 CAPLUS

CN 6,7,8-Isoquinolinetriol, 1-(3,5-dibromo-4-hydroxyphenyl)-1,2,3,4-tetrahydro- (CA INDEX NAME)



OSC.G 14 THERE ARE 14 CAPLUS RECORDS THAT CITE THIS RECORD (14 CITINGS)
 RE.CNT 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN
 AN 2005:409312 CAPLUS [Full-text](#)
 DN 142:441893
 TI Neuromuscular blocking agents and antagonists thereof
 IN Savarese, John J.
 PA Cornell Research Foundation, Inc., USA
 SO PCT Int. Appl., 44 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

PATENT NO.

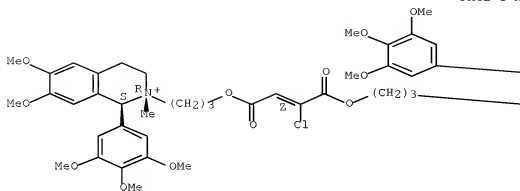
KIND DATE

APPLICATION NO.

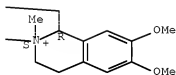
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PI WO 2005041960 A2 20050512 WO 2004-US35869 20041028
 WO 2005041960 A3 20050707
 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
 RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
 US 20050192243 A1 20050901 US 2004-975197 20041028
 EP 1684753 A2 20060802 EP 2004-810087 20041028
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR
 PRAI US 2003-515048P P 20031028
 WO 2004-US35869 W 20041028
 ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT
 OS MARPAT 142:441893
 AB The invention provides methods, compns. and kits for controlling the maximum clin. duration of an ultrashort to intermediate halofumarate neuromuscular blockers. In one embodiment, the methods of the invention involve fast-acting agents that antagonize the neuromuscular blockade caused by administration of a halofumarate neuromuscular blocking agent. Agents that can antagonize the neuromuscular blockade caused by administration of a halofumarate neuromuscular blocking agent include cysteine, N-acetylcysteine, glutathione, as well as related cysteine analogs and combinations thereof.
 IT 213998-46-0, GW 280430A
 RL: ADV (Adverse effect, including toxicity); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (halofumarate neuromuscular blocking agents and antagonists thereof such as cysteine analogs)
 RN 213998-46-0 CAPLUS
 CN Isoquinolinium, 2-[3-[[[(2Z)-2-chloro-1,4-dioxo-4-[3-[(1S,2R)-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-(3,4,5-trimethoxyphenyl)isoquinolinio]propoxy]-2-butenyl]oxy]propyl]-1,2,3,4-tetrahydro-6,7-dimethoxy-2-methyl-1-[(3,4,5-trimethoxyphenyl)methyl]-, chloride (1:2), (1R,2S)- (CA INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.



● 2 Cl-



OSC.G 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD (1 CITINGS)

L6 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2005:284144 CAPLUS Full-text

DN 142:355176

TI Preparation of 6,8-dimethoxyisoquinolines as novel potassium channels modulators

IN Garcia, Gabriel; Saeb, Wael; Kramer, Bernd

PA 4SC AG, Germany

SO U.S. Pat. Appl. Publ., 54 pp.

CODEN: USXXCO

DT Patent

LA English

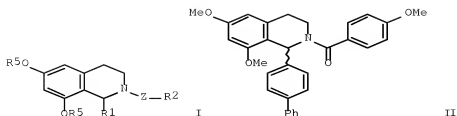
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|-----------------|------|----------|-----------------|----------|
| PI | US 20050070570 | A1 | 20050331 | US 2004-869914 | 20040618 |
| FRAI | US 2003-479159P | P | 20030618 | | |

ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

OS MARPAT 142:355176

GI



AB The invention relates to compds. I [Z = carbonyl, thiocarbonyl or sulfonyl; R1 = alkyl, alkenyl, alkynyl, aryl, H, halo, etc.; R2 = H, OH, CH2SO2alkyl, CH2SO2cycloalkyl, etc.; R5 = alkyl, alkenyl or alkynyl] which are useful for the prevention, alleviation or treatment of diseases, conditions or disorders which are associated with, or dependent on the membrane potential or conductance of cells in mammals, including a human. The general methods for synthesis of compds. I are described. One hundred sixty five compds. I (such as II) were prepared Biol. data were given for representative compds. I. The pharmaceutical composition comprising the compound I is claimed.

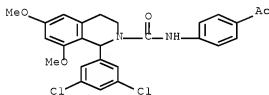
IT 648901-37-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of 6,8-dimethoxyisoquinolines as novel potassium channels modulators)

RN 648901-37-1 CAPLUS

CN 2(1H)-Isoquinolinecarboxamide, N-(4-acetylphenyl)-1-(3,5-dichlorophenyl)-3,4-dihydro-6,8-dimethoxy- (CA INDEX NAME)



OSC.G 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD (3 CITINGS)

L6 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2009 ACS on STN

AN 2005:99335 CAPLUS [Full-text](#)

DN 142:198067

TI Preparation of pyrazolo[3,4-c]isoquinoline derivatives as anaplastic lymphoma kinase modulators

IN Anand, Neel Kumar; Blazey, Charles M.; Bowles, Owen Joseph; Bussenius, Joerg; Costanzo, Simona; Curtis, Jeffry Kimo; Dubenko, Larisa; Kennedy, Abigail R.; Khoury, Richard G.; Kim, Angie I.; Manalo, Jean-Claire L.; Peto, Csaba J.; Rice, Kenneth D.; Tsang, Tsze H.

PA Exelixis, Inc., USA

SO PCT Int. Appl., 239 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.

KIND DATE

APPLICATION NO.

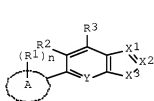
DATE

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|------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----|----------|-----------------|----------|
| PI | WO 2005009389 | A2 | 20050203 | WO 2004-US23762 | 20040723 |
| | WO 2005009389 | A3 | 20050915 | | |
| | W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| | AU 2004259012 | A1 | 20050203 | AU 2004-259012 | 20040723 |
| | CA 2532800 | A1 | 20050203 | CA 2004-2532800 | 20040723 |
| | EP 1648455 | A2 | 20060426 | EP 2004-779018 | 20040723 |
| | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, HR | | | | |
| | JP 2006528638 | T | 20061221 | JP 2006-521273 | 20040723 |
| | US 20070032515 | A1 | 20070208 | US 2006-565657 | 20061005 |
| PRAI | US 2003-489658P | P | 20030723 | | |
| | WO 2004-US23762 | W | 20040723 | | |

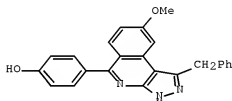
ASSIGNMENT HISTORY FOR US PATENT AVAILABLE IN LSUS DISPLAY FORMAT

OS CASREACT 142:198067; MARPAT 142:198067

GI



I



II

AB The title compds. I [wherein A = (hetero)cycle; R1-R3 = independently H, halo, CN, etc.; X1 and X2 = independently N or (un)substituted CH; X3 = O, S, or (un)substituted NH; n = 1-5; Y = N or (un)substituted CH] or pharmaceutically acceptable salts, stereoisomers, prodrugs, or metabolites thereof are prepared as inhibitors of anaplastic lymphoma kinase (ALK). For example, the compound II was prepared in a multi-step synthesis. Some of compds. I inhibited ALK with IC50 of ≤ 99 nM. I are useful for the treatment of diseases mediated by ALK, including diseases such as cancer, immunol. disorders, cardiovascular diseases, and other degenerative disorders (no data). Formulations containing I as an active ingredient were also described.

IT 938854-82-3P

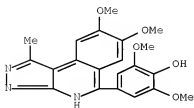
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of pyrazolo[3,4-c]isoquinoline derivs. as anaplastic lymphoma kinase modulators)

RN 838854-82-3 CAPLUS

CN Phenol, 4-(7,8-dimethoxy-1-methyl-3H-pyrazolo[3,4-c]isoquinolin-5-yl)-2,6-

dimethoxy- (CA INDEX NAME)



OSC.G 6 THERE ARE 6 CAPLUS RECORDS THAT CITE THIS RECORD (6 CITINGS)
 RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

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